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(54) Title: GENE EXPRESSION PROFILES IN NORMAL AND CANCER CELLS

(57) Abstract

As a step towards understanding the complex differences between normal and cancer cells, gene expression patterns were examined in gastrointestinal tumors. More than 300,000 transcripts derived from at least 45,000 different genes were analyzed. Although extensive similarity was noted between the expression profiles, more than 500 transcripts that were expressed at significantly different levels in normal and neoplastic cells were identified. These data provide insights into the extent of expression differences underlying malignancy and reveal genes that are useful as diagnostic or prognostic markers.

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Gene Expression Profiles in Normal and Cancer Cells

This invention was made with support from the National Institutes of Health, Grant No. GM07309, CA57345, and CA62924. The U.S. government therefore retains certain rights in the invention.

TECHNICAL FIELD OF THE INVENTION

This invention is related to the diagnosis of cancer, and tools for carrying out such diagnosis.

BACKGROUND OF THE INVENTION

Much of cancer research over the past 50 years has been devoted to the analyses of genes that are expressed differently in tumor cells compared to their normal counterparts. Although hundreds of studies have pointed out differences in the expression of one or a few genes, no comprehensive study of gene expression in the cancer cell has been reported. It is therefore not known how many genes are expressed differentially in tumor versus normal cells, whether the bulk of these differences are cell autonomous rather than being dependent on the tumor microenvironment, and whether most differences are cell-type specific or tumor specific. Thus there is a need in the art for information on the molecular changes that occur in cells during cancer development and progression.

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SUMMARY OF THE INVENTION

According to one embodiment of the invention, a method is provided for diagnosing colon cancer in a sample suspected of being neoplastic. The method comprises the steps of:

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comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 3;

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identifying the first sample as neoplastic when the level of the at least one transcript is found to be lower in the first sample than in the second sample.

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According to another embodiment of the invention, another method is provided for diagnosing colon cancer in a sample suspected of being neoplastic. The method comprises the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2;

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identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

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In another embodiment of the invention an isolated and purified human nucleic acid molecule is provided. The molecule comprises a SAGE tag selected from SEQ ID NO:1-732.

In yet another aspect of the invention an isolated nucleotide probe is provided. The probe comprises at least 12 nucleotides of a human nucleic acid molecule, wherein the human nucleic acid molecule comprises a SAGE tag selected from SEQ ID NO: 1-732.

According to another aspect of the invention a method is provided for diagnosing pancreatic cancer in a sample suspected of being neoplastic. The method comprises the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a pancreatic tissue suspected of being neoplastic and the second sample is of a normal human colon tissue, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4;

identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

According to still another embodiment of the invention a method of diagnosing cancer in a sample suspected of being neoplastic is provided. The method comprises the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a tissue suspected of being neoplastic and the second sample is of a normal human tissue, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 5;

identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

According to another embodiment of the invention a method is provided to aid in the determination of a prognosis for a colon cancer patient. The method comprises the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic colonic tissue and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 3;

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determining a poorer prognosis if the level of the at least one transcript is found to be lower in the first sample than in the second sample.

According to another aspect of the invention a method to aid in determining a prognosis for a patient with colon cancer is provided. The method comprises the steps of:

comparing the level of at least one transcript in a first tissue sample to a second sample, wherein the first sample is of a colonic cancer tissue and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

In yet another embodiment of the invention a method is provided for diagnosing colon cancer in a sample suspected of being neoplastic. The method comprises the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 3;

identifying the first sample as neoplastic when the level of expression of the protein is found to be lower in the first sample than in the second sample.

In another aspect of the invention a method of diagnosing colon cancer in a sample suspected of being neoplastic is provided. The method comprises the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript

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identified by a tag selected from the group consisting of those shown in Table 2,

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

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According to another embodiment of the invention a method is provided to aid in determining a prognosis of a patient having pancreatic cancer. The method comprises the steps of:

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comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic pancreatic tissue and the second sample is of a normal human colon tissue, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4;

determining a poorer prognosis if transcription is found to be higher in the first sample than in the second sample.

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In yet another aspect of the invention a method to aid in providing a prognosis for a cancer patient is provided. The method comprises the steps of:

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comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic tissue and the second sample is of a normal human tissue of the same tissue type, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 5;

determining a poorer prognosis if transcription is found to be higher in the first sample than in the second sample.

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According to still another aspect of the invention, a method is provided for diagnosing pancreatic cancer in a sample suspected of being neoplastic. The method comprises the steps of:

comparing the level of expression of at least one protein encoded by a transcript in a first sample of a tissue to a second sample, wherein the first sample is of a pancreatic tissue suspected of being neoplastic and the second sample is of a normal human colon tissue, wherein said protein is

encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4;

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

According to yet another aspect of the invention a method is provided for diagnosing cancer in a sample suspected of being neoplastic. The method comprises the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a tissue suspected of being neoplastic and the second sample is of a normal human tissue, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 5;

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

In still another embodiment of the invention a method is provided to aid in the determination of a prognosis of a colon cancer patient. The method comprises the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic colonic tissue and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 3;

determining a poorer prognosis if the level of expression is found to be lower in the first sample than in the second sample.

In still another embodiment of the invention a method is provided to aid in determining a prognosis for a patient with colon cancer. The method comprises the steps of:

comparing the level of expression of at least one protein in a first tissue sample to a second sample, wherein the first sample is of a colonic cancer tissue and the second sample is of a normal human colonic tissue, and

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wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 2;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

In still another aspect of the invention a method is provided to aid in determining a prognosis of a patient having pancreatic cancer. The method comprises the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic pancreatic tissue and the second sample is of a normal human colon tissue, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

According to even a further aspect of the invention a method is provided to aid in providing a prognosis for a cancer patient. The method comprises the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic tissue and the second sample is of a normal human tissue of the same tissue type, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 5;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

In still another embodiment of the invention a method of treating a cancer cell is provided. The method comprises the step of:

administering to a cancer cell an antibody which specifically binds to a protein encoded by a transcript identified by a tag selected from the group consisting of those shown in Tables 2, 4, and 5, wherein the antibody is linked to a cytotoxic agent.

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In another aspect of the invention an antibody linked to a cytotoxic agent is provided. The antibody specifically binds to a protein encoded by a transcript identified by a tag selected from the group consisting of those shown in Tables 2, 4, and 5.

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According to another aspect of the invention, a method of detecting colon cancer in a patient is provided. The method comprises the steps of:

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comparing the level of at least one protein or transcript in a first body sample to a second body sample, wherein the first sample is a body sample of the patient and the second sample is of a normal human, wherein the protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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identifying neoplasia when the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

In another aspect of the invention a method of detecting pancreatic cancer in a patient is provided. The method comprises the steps of:

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comparing the level of at least one protein or transcript encoded by a transcript in a first sample of a tissue to a second sample, wherein the first sample is of the patient and the second sample is of a normal human, wherein said protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown Table 4, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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identifying neoplasia when the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

Also provided by the present invention is a method of detecting cancer in a patient. The method comprises the steps of:

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comparing the level of at least one protein or transcript in a first sample to a second sample, wherein the first sample is of patient and the second sample is of a normal human, wherein said protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown Table 5, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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identifying neoplasia when the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

Additionally provided by the present invention is a method to aid in the determination of a prognosis for a colon cancer patient. The method comprises the steps of:

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comparing the level of at least one protein or transcript in a first sample to a second sample, wherein the first sample is of a colon cancer patient and the second sample is of a normal human, wherein the protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown in Table 3, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum:

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determining a poorer prognosis if the level of the at least one protein or transcript is found to be lower in the first sample than in the second sample.

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Provided by another embodiment of the invention is a method to aid in determining a prognosis for a patient with colon cancer. The method comprises the steps of:

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comparing the level of at least one protein or transcript in a first sample to a second sample, wherein the first sample is of a colonic cancer patient and the second sample is of a normal human, wherein the protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

According to still another aspect of the invention, a method to aid in determining a prognosis of a patient having pancreatic cancer is provided. The method comprises the steps of:

comparing the level of at least one protein or transcript in a first sample to a second sample, wherein the first sample is of a pancreatic cancer patient and the second sample is of a normal human, wherein said protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown Table 4, wherein said first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

Also provided by the present invention is a method to aid in providing a prognosis for a cancer patient. The method comprises the steps of:

comparing the level of expression of at least one protein or transcript in a first sample to a second sample, wherein the first sample is of a cancer patient and the second sample is of a normal human, wherein said protein is encoded by a transcript and the transcript is identified by a tag selected from the group consisting of those shown Table 5, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one protein or transcript is found to be higher in the first sample than in the second sample.

The present invention further includes antisense oligonucleotides complementary in whole or in part to SEQ ID NOS:1-732.

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This invention also provides a method for screening for candidate agents that modulate the expression of a polynuleotide selected from the group consisting of the polynucleotides in SEQ ID NOS.1-732 or their respective complements, by contacting a test agent with a pancreatic or colon cell and monitoring expression of the polynucleotide, wherein the test agent which modifies the expression of the polynucleotide is a candidate agent.

The present invention provides the art with new methods and reagents for diagnosing and prognosing cancers. In addition, some of the newly disclosed genes may play an important role in the development of cancers.

BRIEF DESCRIPTION OF THE DRAWINGS

Fig. 1. Comparison of expression patterns in colorectal cancers and normal colon epithelium. (FIG. 1A) A semi-logarithmic plot reveals 51 tags that were decreased more than 10 fold in primary CR cancer cells whereas 32 tags were increased more than 10 fold. 62,168 and 60,878 tags derived from normal colon epithelium and primary CR cancers, respectively, were used for this analysis. The relative expression of each transcript was determined by dividing the number of tags observed in tumor and normal tissue as indicated. To avoid division by 0, a tag value of 1 was used for any tag that was not detectable in one of the samples. These ratios were then rounded to the nearest integer and their distribution plotted on the abscissa. The number of genes displaying each ratio was plotted on the ordinate. Tu: CR tumors; NC: Normal colon. (FIG. 1B and FIG. 1C) Differentially expressed genes in The number of transcripts found to be differentially colorectal cancers. expressed (P < 0.01) are presented as Venn diagrams. Diagrams of transcripts that were decreased (FIG. 1B) or increased (FIG. 1C) in CR cancers compared to normal colon epithelium. Comparisons were between primary tumors and cells in culture as indicated.

Fig. 2. Northern blot analysis of genes differentially expressed in gastrointestinal neoplasia. Northern blot analysis was performed on total RNA (5 µg isolated from primary CR carcinomas (T) and matching normal colon epithelium (N), or pancreatic carcinomas. The top panel in each case show an

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example of the ethidium bromide stained gels prior to transfer. The number of SAGE tags observed in the original analysis is indicated to the right of each blot. (FIG. 2A) Examples of transcripts that were decreased or increased in CR cancers. (FIG.2B) Examples of transcripts increased in pancreatic cancers (10). (FIG.2C) Examples of transcripts elevated in cancer which were or were not cancer type specific. Probes used for Northern blot analysis were as follows (Human SAGE Tag unique identifier, gene name, (GenBank accession number)): (FIG. 2A) H204104, Guanylin (M95714); H259108, (see Table 2); H1000193, (see Table 2); H998030, (see Table 2). (FIG. 2B) H294155, RIG-E (U42376); H560056, TIMP-1 (S68252). (FIG. 2C) H802810, EST338411 (W52120); H85882, 1-8D (X57351); H618841, GA733-1 (X13425).

Tables 2-5. Transcripts Differentially Expressed in Human Cancer.

Tag sequence represents the NlaIII site plus the adjacent 11 bp SAGE tag. Tag number indicates a SAGE UID (unique identifier). NC, TU, CL, PT, PC, refers to the number of the indicated tag observed in RNA isolated from normal colorectal epithelium, primary colorectal cancers, colorectal cancer cell lines, primary pancreatic cancers, or pancreatic cancer cell lines, respectively. The Accession and Gene Name refer to representative GenBank entries that contain the tag sequence.

Table 2 Transcripts increased in colorectal cancer.

Table 3 Transcripts decreased in colorectal cancer.

Table 4 Transcripts increased in pancreatic cancer.

Table 5 Transcripts increased in pancreatic and colorectal cancer.

25 **DETAILED DESCRIPTION**

The inventors have discovered sets of human genes which are either upregulated or downregulated in cancer cells, as compared to normal cells. Specifically, certain genes have been found to be upregulated or downregulated in colorectal and/or pancreatic cancer cells, when compared to normal colon

cells. These sets of differentially regulated genes can be used as diagnostic markers, either individually or in sets of, for example, 2, 5, 10, 20, or 30.

Genes whose expression was detected to be increased in colorectal cancer are shown in Table 2. Genes whose expression was detected to be decreased in colorectal cancer are shown in Table 3. Genes whose expression was detected as increased in pancreatic cancer are shown in Table 4. Genes whose expression was detected as increased in both pancreatic cancer and colorectal cancer are shown in Table 5. These latter genes likely play a role in neoplastic development generally.

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Tag sequences, as provided herein, uniquely identify genes. This is due to their length, and their specific location (3') in a gene from which they are drawn. The full length genes can be identified by matching the tag to a gene data base member, or by using the tag sequences as probes to physically isolate previously unidentified genes from cDNA libraries. The methods by which genes are isolated from libraries using DNA probes are well known in the art. See, for example, Veculescu et al., Science 270: 484 (1995), and Sambrook et al. (1989), MOLECULAR CLONING: A LABORATORY MANUAL, 2nd ed. (Cold Spring Harbor Press, Cold Spring Harbor, New York). Once a gene or transcript has been identified, either by matching to a data base entry, or by physically hybridizing to a cDNA molecule, the position of the hybridizing or matching region in the transcript can be determined. If the tag sequence is not in the 3' end, immediately adjacent to the restriction enzyme used to generate the SAGE tags, then a spurious match may have been made. Confirmation of the identity of a SAGE tag can be made by comparing transcription levels of the tag to that of the identified gene in certain cell types.

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In addition to the sequences shown in SEQ ID NOS: 1-732, or their complements, this invention also provides the anti-sense polynucleotide stand, e.g. antisense RNA to these sequences or their complements. One can obtain an antisense RNA using the sequences provided in SEQ ID NOS: 1-732 and the methodology described in Vander Krol et al. (1988) BioTechniques 6:958.

The invention also encompasses polynucleotides which differ from that of the polynucleotides described above, but which produce the same phenotypic effect, such as the allele. These altered, but phenotypically equivalent polynucleotides are referred to "equivalent nucleic acids." This invention also encompasses polynucleotides characterized by changes in non-coding regions that do not alter the phenotype of the polypeptide produced therefrom when compared to the polynucleotide herein. This invention further encompasses polynucleotides, which hybridize to the polynucleotides of the subject invention under conditions of moderate or high stringency.

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The polynucleotides can be conjugated to a detectable marker, e.g., an enzymatic label or a radioisotope for detection of nucleic acid and/or expression of the gene in a cell. A wide variety of appropriate detectable markers are known in the art, including fluorescent, radioactive, enzymatic or other ligands, such as avidin/biotin, which are capable of giving a detectable signal. In preferred embodiments, one will likely desire to employ a fluorescent label or an enzyme tag, such as urease, alkaline phosphatase or peroxidase, instead of radioactive or other environmental undesirable reagents. In the case of enzyme tags, colorimetric indicator substrates are known which can be employed to provide a means visible to the human eye or spectrophotometrically, to identify specific hybridization with complementary nucleic acid-containing samples. Briefly, this invention further provides a method for detecting a single-stranded polynucleotide identified by SEQ ID NOS.1-732 or its complement, by contacting target single-stranded polynucleotides with a labeled, single-stranded polynucleotide (a probe) which is at least 10 nucleotides of the complement of SEQ ID NOS: 1-732 (or the corresponding complement) under conditions permitting hybridization (preferably moderately stringent hybridization conditions) of complementary single-stranded polynucleotides, or more preferably, under highly stringent hybridization conditions. Hybridized polynucleotide pairs are separated from un-hybridized, single-stranded polynucleotides. The hybridized polynucleotide

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pairs are detected using methods well known to those of skill in the art and set forth, for example, in Sambrook et al. (1989) supra.

The polynucleotides of this invention can be isolated using the technique described in the experimental section or replicated using PCR. The PCR technology is the subject matter of United States Patent Nos.4,683,195. 4.800.159, 4.754,065, and 4.683,202 and described in PCR: The Polymerase Chain Reaction (Mullis et al. eds, Birkhauser Press, Boston (1994)) or MacPherson et al. (1991) and (1994), supra, and references cited therein. Alternatively, one of skill in the art can use the sequences provided herein and a commercial DNA synthesizer to replicate the DNA. Accordingly, this invention also provides a process for obtaining the polynucleotides of this invention by providing the linear sequence of the polynucleotide, nucleotides, appropriate primer molecules, chemicals such as enzymes and instructions for their replication and chemically replicating or linking the nucleotides in the proper orientation to obtain the polynucleotides. In a separate embodiment, these polynucleotides are further isolated. Still further, one of skill in the art can insert the polynucleotide into a suitable replication vector and insert the vector into a suitable host cell (procaryotic or eucaryotic) for replication and amplification. The DNA so amplified can be isolated from the cell by methods well known to those of skill in the art. A process for obtaining polynucleotides by this method is further provided herein as well as the polynucleotides so obtained.

RNA can be obtained by first inserting a DNA polynucleotide into a suitable host cell. The DNA can be inserted by any appropriate method, e.g., by the use of an appropriate gene delivery vector or by electroporation. When the cell replicates and the DNA is transcribed into RNA; the RNA can then be isolated using methods well known to those of skill in the art, for example, as set forth in Sambrook et al. (1989) supra. For instance, mRNA can be isolated using various lytic enzymes or chemical solutions according to the procedures set forth in Sambrook et al. (1989), supra or extracted by nucleic-acid-binding resins following the accompanying instructions provided by manufactures.

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Polynucleotides having at least 10 nucleotides and exhibiting sequence complementarity or homology to SEQ ID NOS: 1-732 find utility as hybridization probes. In some aspects, the full coding sequence of the transcript, i.e., for SEQ ID NOS: 1-732, are known. Accordingly, any portion of the known sequences available in GenBank, or homologous sequences, can be used in the methods of this invention.

It is known in the art that a "perfectly matched" probe is not needed for a specific hybridization. Minor changes in probe sequence achieved by substitution, deletion or insertion of a small number of bases do not affect the hybridization specificity. In general, as much as 20% base-pair mismatch (when optimally aligned) can be tolerated. Preferably, a probe useful for detecting the aforementioned mRNA is at least about 80% identical to the homologous region of comparable size contained in the previously identified sequences identified by SEQ ID NOS:1-732, which correspond to previously characterized genes or SEQ ID NOS:1-732, which correspond to known ESTs. More preferably, the probe is 85% identical to the corresponding gene sequence-after-alignment-of-the-homologous region; even more preferably, it exhibits 90% identity.

These probes can be used in radioassays (e.g. Southern and Northern blot analysis) to detect, prognose, diagnose or monitor various pancreatic or colon cells or tissue containing these cells. The probes also can be attached to a solid support or an array such as a chip for use in high throughput screening assays for the detection of expression of the gene corresponding to one or more polynucleotide(s) of this invention. Accordingly, this invention also provides at least one of the transcripts identified as SEQ ID NOS:1-732, or its complement, attached to a solid support for use in high throughput screens.

The total size of fragment, as well as the size of the complementary stretches, will depend on the intended use or application of the particular nucleic acid segment. Smaller fragments will generally find use in hybridization embodiments, wherein the length of the complementary region may be varied,

such as between about 10 and about 100 nucleotides, or even full length according to the complementary sequences one wishes to detect.

Nucleotide probes having complementary sequences over stretches greater than 10 nucleotides in length are generally preferred, so as to increase stability and selectivity of the hybrid, and thereby improving the specificity of particular hybrid molecules obtained. More preferably, one can design polynucleotides having gene-complementary stretches of more than 50 nucleotides in length, or even longer where desired. Such fragments may be readily prepared by, for example, directly synthesizing the fragment by chemical means, by application of nucleic acid reproduction technology, such as the PCR technology with two priming oligonucleotides as described in U.S. Pat. No. 4,603,102 or by introducing selected sequences into recombinant vectors for recombinant production. A preferred probe is about 50-75 or more preferably, 50-100, nucleotides in length.

The polynucleotides of the present invention can serve as primers for the detection of genes or gene transcripts that are expressed in pancreatic or colon cells. In this context, amplification means any method employing a primer-dependent polymerase capable of replicating a target sequence with reasonable fidelity. Amplification may be carried out by natural or recombinant DNA-polymerases such as T7 DNA polymerase, Klenow fragment of E.coli DNA polymerase, and reverse transcriptase.

A preferred amplification method is PCR. However, PCR conditions used for each reaction are empirically determined. A number of parameters influence the success of a reaction. Among them are annealing temperature and time, extension time, Mg²⁺ ATP concentration, pH, and the relative concentration of primers, templates, and deoxyribonucleotides. After amplification, the resulting DNA fragments can be detected by agarose gel electrophoresis followed by visualization with ethidium bromide staining and ultraviolet illumination.

The invention further provides the isolated polynucleotide operatively linked to a promoter of RNA transcription, as well as other regulatory

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sequences for replication and/or transient or stable expression of the DNA or RNA. As used herein, the term "operatively linked" means positioned in such a manner that the promoter will direct transcription of RNA off the DNA molecule. Examples of such promoters are SP6, T4 and T7. In certain embodiments, cell-specific promoters are used for cell-specific expression of the inserted polynucleotide. Vectors which contain a promoter or a promoter/enhancer, with termination codons and selectable marker sequences, as well as a cloning site into which an inserted piece of DNA can be operatively linked to that promoter are well known in the art and commercially available. For general methodology and cloning strategies, see Gene Expression Technology (Goeddel ed., Academic Press, Inc. (1991)) and references cited therein and Vectors: Essential Data Series (Gacesa and Ramji, eds., John Wiley & Sons, N.Y. (1994)), which contains maps, functional properties, commercial suppliers and a reference to GenEMBL accession numbers for various suitable vectors. Preferable, these vectors are capable of transcribing RNA in vitro or in vivo.

Fragment of the sequences shown in SEQ ID NOS:1-732 or their respective complements also are encompassed by this invention, preferably at least 10 nucleotides and more preferably having at least 18 nucleotides. Larger polynucleotides, e.g., cDNA or genomic DNA, which hybridize under moderate or stringent conditions to the polynucleotide sequences shown in SEQ ID NOS:1-732, or their respective complements, also are encompassed by this invention.

In one embodiment, these fragments are polynucleotides that encode polypeptides or proteins having diagnostic and therapeutic utilities as described herein as well as probes to identify transcripts of the protein which may or may not be present. These nucleic acid fragments can by prepared, for example, by restriction enzyme digestion of the polynucleotide of SEQ ID NOS:1-732, or their complements, and then labeled with a detectable marker. Alternatively, random fragments can be generated using nick translation of the molecule. For

methodology for the preparation and labeling of such fragments, see Sambrook et al., (1989) supra.

Expression vectors containing these nucleic acids are useful to obtain host vector systems to produce proteins and polypeptides. It is implied that these expression vectors must be replicable in the host organisms either as episomes or as an integral part of the chromosomal DNA. Suitable expression vectors include viral vectors, including adenoviruses, adeno-associated viruses, retroviruses, cosmids, etc. Adenoviral vectors are particularly useful for introducing genes into tissues in vivo because of their high levels of expression and efficient transformation of cells both in vitro and in vivo. When a nucleic acid is inserted into a suitable host cell, e.g., a procaryotic or a eucaryotic cell and the host cell replicates, the protein can be recombinantly produced. Suitable host cells will depend on the vector and can include mammalian cells, animal cells, human cells, simian cells, insect cells, yeast cells, and bacterial cells constructed using well known methods. See Sambrook et al. (1989) supra. In addition to the use of viral vector for insertion of exogenous nucleic acid into cells, the nucleic acid can be inserted into the host cell by methods well known in the art such as transformation for bacterial cells; transfection using calcium phosphate precipitation for mammalian cells; or DEAE-dextran; electroporation; or microinjection. See Sambrook et al. (1989) supra for this methodology. Thus, this invention also provides a host cell, e.g. a mammalian cell, an animal cell (rat or mouse), a human cell, or a procaryotic cell such as a bacterial cell, containing a polynucleotide encoding a protein or polypeptide or antibody.

When the vectors are used for gene therapy in vivo or ex vivo, a pharmaceutically acceptable vector is preferred, such as a replication-incompetent retroviral or adenoviral vector. Pharmaceutically acceptable vectors containing the nucleic acids of this invention can be further modified for transient or stable expression of the inserted polynucleotide. As used herein, the term "pharmaceutically acceptable vector" includes, but is not limited to, a vector or delivery vehicle having the ability to selectively target

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and introduce the nucleic acid into dividing cells. An example of such a vector is a "replication-incompetent" vector defined by its inability to produce viral proteins, precluding spread of the vector in the infected host cell. An example of a replication-incompetent retroviral vector is LNL6 (Miller, A.D. et al. (1989) BioTechniques 7:980-990). The methodology of using replication-incompetent retroviruses for retroviral-mediated gene transfer of gene markers is well established (Correll et al. (1989) PNAS USA 86:8912; Bordignon (1989) PNAS USA 86:8912-52; Culver, K. (1991) PNAS USA 88:3155; and Rill, D.R. (1991) Blood 79(10):2694-700. Clinical investigations have shown that there are few or no adverse effects associated with the viral vectors, see Anderson (1992) Science 256:808-13.

Compositions containing the polynucleotides of this invention, in isolated form or contained within a vector or host cell are further provided herein. When these compositions are to be used pharmaceutically, they are combined with a pharmaceutically acceptable carrier.

This invention further encompasses genes, either genomic or cDNA, which code for a polypeptide or protein in the cell of interest. The genes specifically hybridize under moderate or stringent conditions to a polynucleotide identified by SEQ ID NOS: 1-732 or their respective complements. The process of identification of larger fragment or the full-length coding sequence to which the partial sequence depicted in SEQ ID NOS:1-732 hybridizes preferably involves the use of the methods and reagents provided in this invention, either singularly or in combination.

Five methods are disclosed herein which allows one of skill in the art to isolate the gene or cDNA corresponding to the transcripts of the invention.

RACE-PCR Technique

One method to isolate the gene or cDNA which code for a polypeptide or protein and which corresponds to a transcript of this invention, involves the 5'-RACE-PCR technique. In this technique, the poly-A mRNA that contains the coding sequence of particular interest is first identified by hybridization to

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a sequence disclosed herein and then reverse transcribed with a 3'-primer comprising the sequence disclosed herein. The newly synthesized cDNA strand is then tagged with an anchor primer of a known sequence, which preferably contains a convenient cloning restriction site attached at the 5'end. The tagged cDNA is then amplified with the 3'-primer (or a nested primer sharing sequence homology to the internal sequences of the coding region) and the 5'-anchor primer. The amplification may be conducted under conditions of various levels of stringency to optimize the amplification specificity. 5'-RACE-PCR can be readily performed using commercial kits (available from, e.g., BRL Life Technologies Inc, Clotech) according to the manufacturer's instructions.

Identification of known genes or ESTs

In addition, databases exist that reduce the complexity of ESTs by assembling contiguous EST sequences into tentative genes. For example, TIGR has assembled human ESTs into a datable called THC for tentative human consensus sequences. The THC database allows for a more definitive assignment compared to ESTs alone. Software programs exist (give examples) that allow for assembling ESTs into contiguous sequences from any organism.

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Alternatively, mRNA from a sample preparation was used to construct cDNA library in the ZAP Express vector following the procedure described in Velculescu et al. (1997) Science 270:484. The ZAP Express cDNA synthesis kit (Stratagene) was used accordingly to the manufacturer's protocol. Plates containing 250 to 2000 plaques are hybridized as described in Rupert et al. (1988) Mol. Cell. Bio. 8:3104 to oligonucleotide probes with the same conditions previously described for standard probes exxcept that the hybridization temperature is reduced to room temperature. Washes are performed in 6X standard-saline-citrate 0.1% SDS for 30 minutes at room temperature. The probes are labeled with 32P-ATP through use of T4 polynucletoide kinase.

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Table 2 - Transcripts increased in colon cancer

Transcripts increased in only colon primary tumors

compared to normal colon (61 genes)

NC: Normal Colon

TU: Colon Primary Tumor

Ct.: Colon Cancer Cell Line

PT: Pancreatic Primary Tumor PC: Pancreatic Cancer Cell Line

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	H641789	38	144	=	25	13	DS1017	Human fetal brain cDNA 3'-end GEN-00/C04.
LA CATOGCIAGOTITA							D53694	Human fetal brain cDNA 3'-end GEN-117E01.
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15 CATGCCCCGIACAIC	0.000011	2	=	1	2	-	D51021	Human fetal brain cDNA 3'-end GEN-007D07.
16 CATGAGTAGGTGGCC	HIBSOID	2		1			DS1052	Human fetal brain cDNA 3'-end GEN-009C05.
				Ì			D52836	Human fetal brain cDNA 3'-end GEN-089E01.
	U388278	2	124	150	=	23	D83195	Human DNA for Deoxyribonuclease I precursor.
17 CATGCIGIAGICC	01200CH	2	12	78	77	2	DS4113	Human fetal brain cDNA 5'-end GEN-129B05.
18 CATGAGACCCACAAC	H327364	49	101	35	-	6	F15796	H.sapiens mitochondrial EST sequence (102-25) from
19 CATGCATTIOTATA	H874182	78	28	7	0	2		
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21 CATOOCCAACCICCI							D52905	Human fetal brain cDNA 5'-end GEN-091D11.
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22 CATGCCCAICCCII	1100001	12	139	<u>~</u>	×	4	U06452	Human melanoma antigen recognized by T-cells (MAK I
23 CATGTTGGTCAGGCI	0/6/7010	3/2	6	2	2	97		
24 CATGTCCTATIANG	20010011	3	47	1	1-	4	D51004	Human fetal brain cDNA 3'-end GEN-006D02.
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26 CATGATGGCAGGAGI	H461411	-	44	7	~	~		
27 CATGCTANGCGAGG	H713234	-	44	20	2	15	103592	Human ADP/ATP translocase mRNA, 3' end, clone pHA I
28 CATUGOTIOACACT	H97078	0	42	17	100	32	X57352	Human 1-8U gene from interferon-inducible gene fam
29 CATUACCIOIAICCC	C01011H	c	39	0	-	0	H01571	yj33e06.r1 Homo sapiens cDNA clone 150562 5' simil
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							T23865	seq2012 Homo sapiens cDNA clone Cot1374Ft-4HB3MA-3
OLUCIO POPORII	H607576	0	2	-	0	0	M32053	Human H19 RNA gene, complete cds.
33 CATGGCCACCCCIO	H798764	=	12	6	33	2	X67247	H.sapiens rpS8 gene for ribosomal protein S8.
34 CATGTAATAAAGUIU	HR17677		3	~	-	14	T11939	A953F Homo sapiens cDNA clone A953 similar to Mito
35 CATGTACIGCICGGA	1101101							

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2	ראומארראוואר						-	X02490	Human interferon-inducible mRNA (cDNA 1-8).
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5	CATCACCACCATCGC	H165175	0	2	0	0	0		
à	ראומסטמסטיי	2720751	٥	2	-	39	-	103040	Human SPARC/osteonectin mRNA, complete cds.
~	A LOVICATICATICANICACITAL	/ b/ 5 b7 H	٥	2	,	3	,	Ī	
3	TOTTOUTE OF THE CATEGORIES	H310975	0	9	9	7	4	U55217	Human RNA fragment from patients with Cronn's disc
î	ראוסכשווסטוויי	63061311	٥	2	,	~	1		
09	60 ICATGGCCCTCTGCCA	700C10H	2	2	•	:	1	ı	telegraphy NAOTH (COTIN)
:	V. CATOTTACATAACA	H992010	0	2	~	~	9	M94083	Human chaperonin-like protein (m.17.) ilikita, collipiet
٥	CAIGITAGAIAAGA					T		1 27706	Human chaperonin protein (Tcp20) gene complete cds
						1	1	1	

Homo sapiens 18S ribosomal protein (HKE3) mRNA seq. Human mRNA for T-cell cyclophilin. Human DNA for insulin-like growth factor II (IGF-2);

Human Bak mRNA, complete cds.

X07868

0

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73

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8 42

1

28

H379369

CATGCCTAGCTGGAT

U16811

S

4

H482584

CATGCTCCTCACCTG CATGCTTGGGTTTTG

80 6

518912

H.sapiens mRNA for ribosomal protein S18. Human scar protein mRNA, complete cds.

X69150

250

55

8

2

42

H965603

CATGTGGTGTTGAGG

9

CATGTCAGATCTITG

L06432 Y00052

M22146

Transcripts increased in both colon primary tumors and colon cancer

cell lines compared to normal colon (47 genes)

NC. Normal Colon

TU: Colon Primary Tumor

CL: Colon Cancer Cell Line

PT: Pancreatic Primary Tumor

PC: Pancreatic Cancer Cell Line

																						_				
Gene Name	it ibecome a protein 1.28 mRNA, complete cds.	numan noosoniai pioceni ezo mis est	Human mRNA for LLRep3.	H.sapiens BBC1 mRNA	H cariene mRNA for 23 kD highly basic protein	11.3d perior factor 7	Hisapiens mkuk tor cronigation taxos	H.sapiens S19 ribosomal protein mKNA, complete cus	Human acidic ribosomal phosphoprotein P2 mRNA, com	If spriene has mRNA for uracil DNA glycosylase.	In Section 1 - 3 - Learnest dehydrospace mRN	Human glyceraidenyde 3-piiospiiaic deilydiogenase mini	H saniens mRNA for elongation factor- I-gamma.	AND AND THE STATE OF THE STATE	Haman pancreatic tumor-related process singles, 2 cm	H.sapiens mRNA for ribosomal protein L8.	transaction and for elboomal profein 1.3.	n.sapiciis inivity tol mosciniis process	Human novel gene mRNA, compiete cds.	Human Wilm's tumor-related protein (QM) mRNA, comp	laminin recentor homolog (3' region) [human, mRNA		H.sapiens mKNA for OKr.	Human mRNA for ribosomal protein L32	Human ribosomal protein S4 (RPS4X) isoform mRNA, c	
Accesion	444531011	014969	X17206	X64707	V66033	A30734	Z1169Z	M81757	M17887	V62770	1	102642	711531	1	M55409	728407		X/3400	M73791	M64241	035050	20000	X80822	X03342	1	1
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	Tag_Number	H599350	1200633	H237333	H355689	H171113	H148949	100000	H207774	H671654	H807748			H959498			H55227	H660601	11174037	H1/403/			LIAAKRZ	COOLAL	H935680	H861056
	Tag Sequence	TOUCH TOUCH TO	CAIGGLAGCCAICCO	2 CATGATGGCTGGIAI	CATGCCCGTCCGGAA	ACOUTACOUTACO	0,0000000000000000000000000000000000000	SCATGAGLACTICCAG	6 CATGCTGGGTTAATA	7 CATGGGATTTGGCCT	CATCTACCATCAATA	CATOLACCATCATA		JUST A A GOOD A	CATOTOGCCAAAGCC		IN CATGAATCCTGTGGA	CA CITCA COCCE.	11 CATGGGACCACTGAA	CATGAGGCTTCCAA			. 00	13 CATGAAGGIGGAGGA	14 CATGTGCACGTTTTC	15 CATGTCAGATCTITG
	*		_	7	-	<u> </u>	7	~	9	-	1	×		ľ	^		2	2	=	12	L	1		=	<u>-</u>	ΙΞ

D14530 Human homolog of yeast ribosomal protein S28, comp	H.sapiens HRPL4 mRNA.	D23661 Human mRNA for ribosomal protein L37, complete cds		M17886 Human acidic ribosomal phosphoprotein P1 mKNA, com	X63527 H.sapiens mRNA for ribosomal protein L19.	M24194 Human MHC protein homologous to chicken B complex	U14967 Human ribosomal protein L21 mRNA, complete cds.	X55954 Human mRNA for HL23 ribosomal protein homologue.	X52839 Human mRNA for ribosomal protein L17.	П	H71935 ys15f12.r1 Homo sapiens cDNA clone 214895 5.	Z43914 H. sapiens partial cDNA sequence; clone c-10d03.		X04347 Human liver mRNA fragment DNA binding protein UP1	\neg	X61156 [H.sapiens mRNA for laminin-binding protein.	103799 Human colin carcinoma faminin-binding protein mRNA	U02032 Human ribosomal protein L23a mRNA, partial cds.	U14970 Human ribosomal protein SS mRNA, complete cds.		M36981 Human putative NDP kinase (nm23-H2S) mRNA, complet	L16785 Homo sapiens c-myc transcription factor (puf) mRNA	L10376 Human (clone CTG-B33) mRNA sequence.	S80520 CAG-isl 7 (trinucleotide repeat-containing sequenc	M77349 Human transforming growth factor-beta induced gene				\neg	\neg	D28137 [Human mRNA for BST-2, complete cds.	Soares senescent fibroblasts NbHSF Homo sapiens cDNA clone		X72718 H. sapiens DNA for orphan ICR V-beta segment (aller
103	8	ê.	118	66	146	77	19	120		99				&	0	57		18	╀	╁	_	-	<u>=</u>	├	∞	∞	2	22	25	3		-	9	Н
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	\neg	_	22 CAIGCAAIAAAIGII	_	_		\neg	_	28 CAIGAIICICCAGIA	\rightarrow	29 CATGACTCCAAAAAA			COLUMN TANGET	30 CATGOTOTICALISC		32 CATGGAAAAA UU I	_	33 CATGAAGAAGATAGA	14 CATGCCTTCGAGATC	15 CATGACIGGGICIAL			36 CATGCAGCICACIGA	A TOTAL CHOCK	37 CA1001010101A	S (Alcolocociavoc	39 CAIGGITCACAITAG	40 CATCIOAAAAAAA	TICATOTOTOTOTO	42 CA1010C10CC1011		A CA CO CE A CIECCE CO	43 (A10(10A1000A0

									2000FC
									Soares fetal heart NoHHI9W Homo sapiens CUIVA Clone 342320
		1161611	•	12	12 16 5	~	7	H121311	
=		1161211	>	7.	2	1	-	Ī	Transfer of the state of the st
1						_			EST176663 Colon carcinoma (Caco-2) cell line il Homo sapiens
								A A 305589	AA305589 CDNA 5' end
_									
1		71 78 01 71 0 777	٥	2	9	Ca	17		X53416 Human mRNA for actin-binding protein (filamin) (AB
Y	14 ICA TOCOCA AGGACC	H010400	>	7	`	70			
•	20000		ľ		90	63	•	17607	vo2761 Human mRNA for fibronectin (FN precursor).
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cell lines compared to normal colon (181 genes) Transcripts increased in only colon cancer

NC: Normal Colon
TU: Colon Primary Tumor
CL: Colon Cancer Cell Line
PT: Pancreatic Primary Tumor

PC: Pancreatic Cancer Cell Line

Ton Sequence	Tag Number	NC	TU	CL PT	r PC		Gene Name
1 ag Schacher	H978825	٦	79	487 136	6 412	X16869	Human mRNA for clongation factor 1-sipnia
CAIGIOIGIAGO	H615043	12	99	265 105	5 125	X53505	Human ribosomal protein S12.
CATUCCCCACCACC	87.PLYCH	137	83	245 36	502	X12883	Human cytokeratin 18.
CATGCAACCAICCA	9178761	+-	+-	╀—	130	L19739	Homo sapiens metallopanstimulin (MPS1)
CATGCACAAACGUIA	050072	╁	+-	186	120	X83412	H.sapiens B1 mRNA for mucin.
CATGAAAAAAAAA		╫	+-	+-	+-	Z32564	H.sapiens FRGAMMA mRNA (819bp) for folate receptor
		\dagger	+	+	-	X76180	H.sapiens mRNA for lung amiloride sensitive Na+ ch
		\dagger	\dagger	+	+	U08470	Human FR-gamma' mRNA, complete cds.
		+	\dagger	+	-	U08471	Human folate receptor 3 mRNA, complete cds.
O.L.O. G.	37770111	E	28	179	104 358	S64030	Human L41 ribosomal protein
CATGTTGGTCCTCTG	07470011			╄	0 87	T91925	ye02/02,r1 Homo sapiens cDNA clone 116571 5'.
CATGTCTCCATACCC	H906438	-	+	┿	+	┸	H caniens ribosomal protein L37a.
CATGAAGACAGTGGC	1.133979	2	-+-	-+-		┙	Human ribosomal profein \$16
CATGCCGTCCAAGGG	H374027	읾	-+				Human themosin heta 10
CATGGGGAAATCGC	H696375	8	g	130	-	10624M	Italian in most control of the contr
CATGAAGGAGATGGG	H41531	30	37	133 3	38 161	X69181	H.sapiens mkrvA for frousening protein E.S.:
CATCGAGGGAGTTC	11567488	38	53	112 6	65 142		Human ribosomal protein L./ /a
A TOTO TOTO TA	H424694	42	8	Ξ	53 49		H.sapiens ribosomal protein L.I.
CAIGCOCTOTOTOTOTOTOTOTOTOTOTOTOTOTOTOTOTOTO	661819H	8	39	109 2	28 120	103537	Human ribosomal protein S6
CA16000010000	H\$49145	32	85	105	44 70	US8682	Human ribosomal protein S28 mRNA, complete cds
CATGGACGACACGAG	19877367	36	48	-	44 65	X52839	Human mRNA for ribosomal protein L17
CATGICACCCACACC	H416106	82	43	8	52 184	U12465	Human ribosomal protein L35
CATGCGCCCCCGCC	H475448	27	14	╁	27 145	M17885	Human acidic ribosomal phosphoprotein PO
CATOCICACCICIC	H955718	2	8	08	46 55	M23725	Human M2-type pyruvate kinase mRNA, complete cos.
CAIGINGCCCCACCC				-	-	M26252	Human TCB gene encoding cytosolic thyroid normone-
	00103611	2	Q	 	92 145	M11147	Human ferritin L chain
20 CATGCCCTGGGTICI	H335102			┨ .		1	

1 LOTTOR CONTINUE OF THE	H150997	0	0	111	0	H09058	y196f11,r1 Homo sapiens cDNA clone 45943 5'.
CATUAUCATCICCAU			T	T	-	Z44640	H. sapiens partial cDNA sequence; clone c-26b05.
			T		-	N75111	yz29e01.r1 Homo sapiens cDNA clone 284472 5'.
O TOTOTOTOTO	H621369	24	32	12	33 99	M31520	Human ribosomal protein S24 mRNA.
CATGOCCIOINIONO	H161624	8	8	26	21 67	777ESX	Human L23 mRNA for putative ribosomal protein.
21222121200101					_		gb AA223340 AA223340 Homo sapiens cDNA clone 650651 5 Similar to
CATGCCAGGAGGAAT	H338081	27	12	-	23 87		gb: Y00371 mai HEAT SHOCK COGNATE 71 KD PKOTEIN (HUMAN)
CATGGGCAAGCCCCA	H672342	8	55	72	27 61		Human Csa-19
CATGAGGAAAGCTGC	H163999	<u>-</u>	42	70	32 146		H.sapiens EST sequence (135-18) from skeletal muscle
CATGAGGGGGAA	H26261	53	46	69	54 79	_	Homo sapiens macrophage migration innibitory factor
CATOCAGAGAGAG	H335945	23	39	99	42 148		H.sapiens ribosomal protein L30.
CATUCCAGACAGAC	H615736	6	2	85	10 22	USS017	Human transketolase (TKT)
CAIGUCCUCCAICIC	H769045	92	<u>e</u>	23	17 76	L25899	Human ribosomal protein L10
CATGOTOTACCAG	H383489	6	=	8	23 46	226876	H.sapiens ribosomal protein L38.
CATOCCICOGRAM	H177610	2	27	2	43 41	X06547	Human class Pi glutathione S-transferase
CATGAGGICCIAGCC	H775658	=	78	8	32 96	X65923	H.sapiens fau mRNA.
CATGGIICCIIGGCC	H796831	3	85	╁	42 68	07777X	H.sapiens RPS26
CATGIAAGGAGCIGA	178673	-	4	╁	┼	WS2460	zc45ei I.rl Soares senescent fibroblasts NbHSF Homo
CAIGAACIAAAAAA				T	-	N92893	2b71h03.s1 Homo sapiens cDNA clone 309077 3'.
O V J J J J J J J J J J J J J J J J J J	H260949	-2	=	57	6	X14957	Human hmgl mRNA for high mobility group protein I.
CATGATTOTCCAO	H200576	=	27	2	30 69	U14973	Human ribosomal protein S29
CATGATAATICITIO	97282H	~	12	12	5 85	U14990	Human XP1PO ribosomal protein S3 (rpS3)
CATGCCCCAGCCAGI	05/24511	1	=	8	-	1 L11566	Homo sapiens ribosomal protein L18 (RPL18)
CATGGGAGIGGACAL	H786433	: =	∞	8	╌	5 H08238	y187a01.r1 Homo sapiens cDNA clone 44932 5'.
CAIGIAAAAAAAAA	H769605	6	71	84	21 47	_	H.sapiens ribosomal protein S13.
CATGGTGTTGCACAA	HK08595	و	7	47	15	5 U31657	Human unknown protein mRNA, partial cds.
כעומתרשמרהרשמה					-	H41030	yn92a10.r1 Homo sapiens cDNA clone 175866 5.
	HK85384	14	22	12	23 15	5 M16660	Human 90-kDa heat-shock protein
CATGGGCICCCACIO	HRANGRA	6	0	8	2	NS7419	yw82e04.rl Homo sapiens cDNA clone 258750 5' simil
CATGLCAACTICTOO	H583573	9	12	9	-	18 X59357	Human mRNA for Epstein-Barr virus small RNAs (EBER)
CATGGATGCTGCCAA					-	L21756	Homo sapiens acute myeloid leukemia associated protein
					\vdash	D17652	Human mRNA for HBp15/L22, complete cds.
CATOAATAGGTCAA	H51925	=	=	46	47 5	S3 M64716	Human ribosomal protein S25
CATCCCTTTAAGGA	H655115	∞	97	45	22 63		Homo sapiens ribosomal protein S20 (RPS20)
CATGOCITITAGOS	H58511	7	12	44	6 2	27 M61831	Human S-adenosylhomocysteine hydrolase (AHCY)
48 CATGAAIGCAGCAG	110011	1		1			

221507 Human elongation factor 1 delta (EF 1delta)	M13932 Human ribosomal protein SI7 mRNA	M10036 Human triosephosphate isomerase		L 19527 Homo sapiens ribosomal protein L27 (RPL27)	X63237 H.sapiens Uba80 mRNA for ubiquitin.	Unknown	X69391 H.sapiens ribosomal protein L6.	H11182 ym14a02.r1 Homo sapiens cDNA clone 47866 5'	T40302 ya31g04.r5 Homo sapiens cDNA clone 62262 5'	T89480 yd98a05.rl Homo sapiens cDNA clone 116240 5'	H01362 yi99c06.rl Homo sapiens cDNA clone 147370 5'		T49412 ya75b09.r1 Homo sapiens cDNA clone 67481 5'.	X07270 Human heat shock protein hsp86.	M91670 Human ubiquitin carrier protein (E2-EPF)	X74070 H.sapiens transcription factor BTF 3.	V00599 Human beta-tubulin	X84694 H.sapiens mRNA for elongations factor Tu-mitochondria	L38995 Homo sapiens nuclear-encoded mitochondrial elongatation factor	S75463 P43=mitochondrial elongation factor homolog (liuman	H48893 yq80b12.r1 Homo sapiens cDNA clone 202079 5'	X71973 H.sapiens GPx-4 mRNA for phospholipid hydroperoxidase	M95787 Human 22kDa smooth muscle protein (SM22)		R74294 yi57f06.r1 Homo sapiens cDNA clone 143363 5.		F17005 H.sapiens EST sequence (011-T1-18) from skeletal muscle	H10519 yl90g04.r1 Homo sapiens cDNA clone 45563 5'.		\neg		1	X52317 Human histone H2A.Z.
22 2	18 N	42 N	49 F	87 1	51	6	42	25			2	14	_	25	1	_	40	=			9	107	0	25		=	98	2	0	26	24	46	9
0	∞	15	35	2	4	0	4	12		\dagger	32	<u>_</u>		4	╌	∞	22	2	T	T	0	<u>∞</u>	Z	7		٥	6	-	0	25	∞	1	7
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<u>®</u>	9	56	24	-	4	0	=	-	T	1	2	2	\vdash	2	0	∞	=	9		T	4	~	5	2		~	87	2	9	6	7	28	4
8	0	4	4	∞	9	0	1	∞			=	3		E	7	=	7	0			12	2	0	~		E	2	-	0	∞	2	6	_
H610939	H678334	H928269	H968173	H672265	H28737	H837237	H803369	H770486			H558943	H217399		CC\$915H	H501287	H493633	H74951	H602783			H119302	H621035	H76231	H528067		H533798	H988366	H1023249	H874103	H246019	H298495	H777109	H552683
49 Transcream	Т	SI CATGAGGGAATAA	_	7		_	_	Т	יי כאוסטוואארטורככ		CATCCAGACTCCTGC	SO CATGATCACACACGC	_	_	60 CATGGAAGCTTGCA	_	Т	6) CATGRACONCETED!	SA CALGOCA MOCCIOS		& OO & OTTOO TANK	6) CATOCATOLICATORY	\neg	\neg	200000000000000000000000000000000000000	ADDA ODDA ODDA ODDA	20 CATGTTACCATATCA		7	Т	\neg	$\neg \neg$	$\overline{}$

	63503711	-		77	0	~	M33680	Human 26-kDa cell surface protein TAPA-1
\neg	H436/33	-	• -	+	┿	<u>≃</u>	Т	Homo sapiens dbpB-like protein
$\overline{}$	H /04300	-	- 0	15	╁╴	2 2	T	Human translational initiation factor 2 beta subunit
$\overline{}$	11504051	- 14	0	1/2	-	29	W07137	za92a11.r1 Soares fetal lung NbHL 19W Homo sapiens
80 CATGGCACAAGAAGA	1004601	,	1	+	╁╴	-	D20503	Human HL60 3'directed Mbol cDNA, HUMGS01477, clone
			1		\dagger		N91592	Soares fetal lung NbHL19W Homo sapiens cDNA clone 303055 3:
		T	T	T	\dagger	-		yv84c07.s1 Homo sapiens cDNA clone 249420 3' similar to contains Alu
							H83884	repetitive element:
7	H908373	1	=	92	=	13	222572	H.sapiens CDEI binding protein mRNA.
81 CATOLCIACCAC					-	-	L09209	Homo sapiens amyloid protein homologue mRNA, compl
		T	T		\vdash	-	L19597	Human binding protein mRNA, partial cds.
		T	T		\vdash	\vdash	86009S	APPH=amyloid precursor protein homolog [human, pla
$\neg \uparrow$	H783697	-	0	22	6	6	W07587	zb06f02.rl Soares fetal lung NbHL19W Homo sapiens
82 CATOUTTICCCAAG		Τ	T	T		-	N28502	yx36f06.r1 Homo sapiens cDNA clone 263843 5
		T	T		\vdash	┝	N35630	yx62a03.rl Homo sapiens cDNA clone 266284 5'
	9CP88LH	2	-	25	6	2	240265	H. sapiens partial cDNA sequence; clone c-1xe03.
8) CATGCCTOTCCAGCC	27.00011	T	1			\vdash	W02723	zc65c03.s1 Soares fetal heart NbHH19W Homo sapiens
			T	T	-	\vdash	N24893	yx99h09.s1 Homo sapiens cDNA clone 269921 3'.
			T	T	\vdash	\vdash	N32178	yy25b09.s1 Homo sapiens cDNA clone 272249 3'.
	H865503	~	2	22	5	-	H21873	y134b10.s1 Homo sapiens cDNA clone 160123 3' simil
84 CATGICALCAICION	50550011			1	\vdash		H26394	yl48e12.s1 Homo sapiens cDNA clone 161518 3' simil
				T		\vdash	H69857	yr88d02.s1 Homo sapiens cDNA clone 212355 3' simil
				1	\vdash	-	H70714	yu69b11.s1 Homo sapiens cDNA clone 239037 3' simil
\neg	H158783	~	∞	22	9	E	X55110	Human mRNA for neurite outgrowth-promoting protein
\neg	H617048	-	-	24	6	 -	X03168	Human mRNA for S-protein.
% CATGOCCOOCCO					T			2032d09.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 588593
	H1023233	7	-	24	7	7	AA143561	3' similar to contains LTR7.t1 LTR7 repetitive element
87 CAIGIIGCICAAAAA	2000				\vdash	├-		2001g11.51 Stratagene colon (#937204) Homo sapiens cDNA clone 566468
							AA152342	3' similar to contains LTR7.13 LTR7 repetitive element;
						\vdash		2186h I 1.51 Stratagene colon (#937204) Homo sapiens cDNA clone 511557
							AA115727	3' similar to contains LTR7.t1 LTR7 repetitive element
	H262987	9	7	24	5	2	R76502	yi6109.r1 Homo sapiens cDNA clone 143753 5.
88 CATOCAAAATCAGGA						-	T32681	EST52915 Homo sapiens cDNA 5' end similar to None.
		\perp			Г		T34662	EST72468 Homo sapiens cDNA 5' end similar to None.
	H533435	上	5	23	4	7	H04634	lyj49h03.rl Homo sapiens cDNA clone 152117 5.
89 CATGGAAGATGTGGG								

				T	r		\vdash	F00364	H. sapiens partial cDNA sequence; clone 76D12; ver
5	CATCCTCATTCA	H761150	0	∞	23	9	4	H01503	yj21c05.s1 Homo sapiens cDNA clone 149384 3'.
3	ראוומחומרוכאווכא					\vdash	-	H84813	yv86c02.s1 Homo sapiens cDNA clone 249602 3' simil
						-		H84956	yv88f07.s1 Homo sapiens cDNA clone 249829 3' simil
ā	CATGGCTTTACTTTG	H654464	4	~	23	6	5	L38961	Homo sapiens putative transmembrane protein (BS)
- 6	CATGTTTCTGAAA	H1046401	9	=	23	<u>_</u>	0.	104026	Human thioredoxin (TXN) mRNA
; 5	CATGTTGCTCACACA	H1023250	-	4	22	0	4	D11078	Human RGH2 gene.
70	CATGGATTTCTCAGC	H589267	0	0	22	0	61		Human mRNA for placental-like alkaline phosphatase
S	CATGAGGAGGGAGGC	H166539	2	3	22	2	4	\exists	Human pyrroline 5-carboxylate reductase mRNA,
:\\	CATGCTTAACCTGG	H651359	3	4	22	7	4	П	Human glutamate dehydrogenase
3 5	CATCCTCTTCGAGAA	H490889	4	∞	22	27	6	П	Human mRNA for glutathione peroxidase
8	CATGAGAACAAAACC	H132098	-	7	21	\dashv	9	-1	H.sapiens mRNA for proliferation-associated gene
g	CATGCCCAGGGAGAA	H346761	3	٣	21	7	24	\Box	Human stimulator of TAK KNA binding (SKB)
:								D16933	Human HepG2 3' region cDNA, clone hmd4111.
10	CATGCACTTCAAGGG	H294155	0	3	2	47	107	U42376	Human retinoic acid induced RIG-E
3		H631331	7	3	2	4	_		Unknown
3 3	CATGTTACCTCCITC	11989024	4	7	20	3	77	F17524	H.sapiens EST sequence (012-12-32) from skeletal in
3 3	CATGACTCTGCCAAG	H122449	4	7	20	5	7	\neg	Unknown
3 3	UT CATIGTOR GATGGCGT	H861095	-	9	61	2	7	W52942	zc03h05.rl Soares parathyroid tumor NoFil/A Homo sap
ž	CATGGGGCTTTTTT	11679936	-	3	19	S	~		yg48h11,r1 Homo sapiens cDNA clone 35917 3 simila
٤	CATGINGACGCCTG	H951912	0	0	61	0	0	\Box	Human lipoprotein apoA1.
3 3	0.1000.00.00.00.00.00.00.00.00.00.00.00.	H386904	0	~	61	9	5	M80244	Human E16 mRNA
2 3	CATGGGGACACCCA(C)	1-1607318	7	9	<u>8</u>	81	15		y158c11.s1 Homo sapiens cDNA clone 162452 3' simil
2		H249854	7	3	18	5	20		H.sapiens ribosomal protein L7.
		H529899	2	-	82	5	15		EST12509 Uterus tumor 1 Homo sapiens cDNA 5' end
2 =		H686319	-	5	81	∞	1	U09510	Human glycyl-tRNA synthetase
=		H855049	3	10	∞	4	4	X76013	H. sapiens QRSHs mRNA for glutaminyl-tRNA synthetas
: =		H11785	0	7	11	0	~	W16529	zb10a11.rl Soares fetal lung NbHL19W Homo sapiens
								W35192	zc70b05.rl Soares fetal heart NbHH19W Homo sapiens
								W52451	zc45d09.rl Soares senescent fibroblasts NbHSF Homo
	CATGCACGCGCTCAA	H288373	0	-	17	0	3	D38251	Human mRNA for RPB5 (XAP4)
		H28872	-	9	17	13	31	D52570	Human fetal brain cDNA 5'-end GEN-081G12.
≘								D52758	Human fetal brain cDNA 5'-end GEN-087A08.
								D55953	Human fetal brain cDNA 5'-end GEN-407H12.
1	116 CATGETGTACCTGGA	H504187	_	0	17.	12	9	M22490	Human bone morphogenetic protein-2B (BMP-2B)
2									

13 CATOCOACOCACGC	H398663	7	9	_	20 -	_ >	M12529	Human apolipoprotein E
CATGLAGAAAAAAAA	H819213	0	-	2	7	-	X16539	H.sapiens RNA for neuroleukin gene.
			T	Γ	-	┞	M27691	Human transactivator protein (CREB) mRNA, complete
CATCATCTTGAAAGG	H228867	0	0	9	2	3	M86667	H.sapiens NAP (nucleosome assembly protein)
CATGGGGGGGGT	H302741	0	-	9	14	0	X53743	H.sapiens mRNA for fibulin-1 C.
CATGATCTTGAAAGG	H228867	0	0	2	~		226328	H. sapiens partial cDNA sequence; clone HEC059
CATGATCTTGAAAGG	H228867	0	0	2	~	~	Z26328	H. sapiens partial cDNA sequence; clane HEC059
CATGGTGGAGGTGCG	H762554	2	2	91	3	5	U22055	Human 100 kDa coactivator mRNA
CATGGTGGACCCCAA	H762197	E	~	2	7	01	R91724	yp98e02.r1 Homo sapiens cDNA clone 195482 5' simil
					\vdash	┝	WS1770	zc48a02.rl Soares senescent fibroblasts NbHSF Homo
				T	\vdash		N42086	Jyy05b03.r1 Homo sapiens cDNA clone 270317 5'
CATCGAGCAGCTGGA	H561787	0	~	2	7	4	R80990	yi94c02.r1 Homo sapiens cDNA clone 146882 5'
						-	R95056	yq44f01.r1 Homo sapiens cDNA clone 198649 S' simil
TOUGHTOUGHT	H633002	-	9	2	∞	7	F16507	H.sapiens EST sequence (147-09) from skeletal musc
200000000000000000000000000000000000000				T	\vdash	-	T50201	yb77h05.r1 Homo sapiens cDNA clone 77241 S' simila
CATCATTCCCTTAAA	H256497	-	∞	2	0	9	S85655	Human prohibitin
CATCCAAAATTTAA	H524541	0	-	2	4	0	M38188	Human unknown protein from clone pHGR74 mRNA, comp
CATGGATCACAGTTT	H577840	0	~	2	0	0	Y00711	Human lactate dehydrogenase B (LDH-B).
CATGAGCCTTTGTTG	H155632	E	7	2	23	5	D83174	Human collagen binding protein 2.
CATGTCTCACCTCC	H910430	0	0	2	0	2	X70940	H.sapiens clongation factor 1 alpha-2.
CATGAGGGGGG	H18469	0	7	2	9	=	T30623	EST19638 Homo sapiens cDNA 5' end similar to None.
						-		HUMGS0004747, Human Gene Signature, 3'-directed cDNA
				···			C01011	sequence.
					╁	╁		zm62d06.s1 Stratagene fibroblast (#937212) Homo sapiens cDNA clone
							AA111865	
				T	\vdash	\vdash	W56516	2d16c08.r1 Soares fetal heart NbHH19W Homo sapiens
CATCTCTTCAGGACC	H980130	-	-	4	~	=	H30299	yo77d04.r1 Homo sapiens cDNA clone 183943 5' simil
2000						-	H50265	yo28c02.rl Homo sapiens cDNA clone 179234 5'.
CATCTAGATAATGGC	H822331	E	4	4	9	4	W01702	za37a06.rl Soares fetal liver spleen INFLS Homo sa
					-	\vdash	W04495	za58b10.r1 Soares fetal liver spleen INFLS Homo sa
				Γ	-		W23528	zc71g11.s1 Soares fetal heart NbHH19W Homo sapiens
CATGCTTAATCCTGA	H508767	0	9	4	9	12	DI 1838	Human HepG2 3'-directed Mbol cDNA, clone hm02e09.
CATOCICAGAGGACC	H673954	0	9	14	5	=	X75598	H.sapiens nm23H1 gene.
CATCTGACTGAGGC	H925194	0	2	4	-	0	T35470	EST85850 Homo sapiens cDNA 5' end similar to None.
22000120010		I	T		t		700000	Portococi tions senions of NA Chand cimilar to None

T35545 EST87066 Homo sapiens cDNA 5' end similar to None.	H01694 yj33g11.s1 Homo sapiens cDNA clone 150596 3.	N78851 zb17d08.s1 Homo sapiens cDNA clone 302319 3'.	N78931 za92h06.s1 Homo sapiens cDNA clone 300059 3'.	H90469 yv01e06.r1 Homo sapiens cDNA clone 241474 5' simil	R76765 yi63g01.r1 Homo sapiens cDNA clone 143952 5' simil	T35045 EST79335 Homo sapiens cDNA similar to None.	H51447 Joo31a05.rl Homo sapiens cDNA clone 179504 5.	W46469 zc32c05.rl Soares senescent fibroblasts NbHSF Homo	W51800 zc48e04.rl Soares senescent fibrobiasts NbHSF Homo	R33196 Jyh77f08.r1 Homo sapiens cDNA clone 135783 5'.	J04799 Human prothymosin-alpha	D80012 Human KIAA0190 protein	U02389 Human hLON ATP-dependent protease mRNA	729819 EST96617 Homo sapiens cDNA 5' end similar to ATP-d	X14850 Human histone H2A.X.	J04088 Human DNA topoisomerase II (top2) mRNA	K01891 Human beta globin retrovirus-like repetitive element	1188396 ESTZ8e05 Homo sapiens cDNA clone 28c05	X74796 H.sapiens p85Mcm mRNA.	D28480 Human mRNA for hMCM2, complete cds.	D55716 Human B lymphoma mRNA for P1cdc47, complete cds.	T30327 EST14849 Homo sapiens cDNA 5' end similar to None.	T34394 EST66942 Homo sapiens cDNA 5' end similar to None.	T47475 yb14c03.rl Homo sapiens cDNA clone 71140 5'.	T50289 yb14h08.rl Homo sapiens cDNA clone 71199 5.	Unknown				\neg	Z49216 H.sapiens mitoxantrone-resistance associated mkNA.	Unknown	\neg	M93651 Human set gene
\vdash	-		-	=	\vdash	\vdash	6		\vdash		2	12	~		9	7	0		∞		-	=		-	T	-	7	4	12	0	4		1	∞
	7			0			2				∞	∞	7		-	 -	0		-			9				2	2	4	4	0	~		٥	1
	4			13			2				13	2	5		13	2	=		=			=				=	=	12	12	12	12	12	12	
	-			4			9				2	~	~		~	·	0		2			~			\perp	上	2	9	S	0	2	-	0	7
	0	_		<u> -</u>	_	1	0	1		$oldsymbol{\perp}$	上	0	-		<u> </u> -	· c	0	1	<u> </u> -	·	1	10	1	1	\downarrow	0	0	0	0	0	0	0	0	
	H576495			H765573			H961304				H1003313		H125315		30P9CSH	57797CH	H16303		H496114			H\$3120	121001			H890535	H697495	H329737	H1048113	H977034	H345789	H63325	H548203	H921067
	23 CATGGATAGTTGTGG	_		24 TOCTOCTOCACAC	200000000000000000000000000000000000000			יייייייייייייייייייייייייייייייייייייי			A CATCATTATATA	CATCUTCATCTCTCTCTAC(T)	TO A COLUMNIA OF COLUMNIA	142 CATGACTOCCOARG	VOLOUVOT TOOL T	145 CATGGAAAGAGCTGA		LAS CATGANATITICATOR	$\overline{}$	146 CATOCTOCACTIACT		_	147 CATGAATATTGAGAA			JUJUJUJUJUTUT VI		SO CATOCCAAGAAGAA	151 CATGTTTTGATAA		CATGCCACGGTTAG	134 CATGAATTCTCCTAA	SS CATGRACCTCCGGGC	156 CATGTGAATCTGGGT

H843485 0			H884181	6	~	=	4	∞	X15804	Human alpha-actinin.
CATGGCGACTCCCCAACT CATGGCATCCCTCCAACT CATGGCATTCCTCCA H141444 0 0 11 1 17 236249 CATGGCAATTCCTCCAA H540023 0 3 11 3 1 180776 AA02809 CATGGCAACTCCTCCAACT H550274 0 1 1 1 0 AA02809 CATGGCACACCACACT H550274 CATGGCAACACACACACACACT H656453 CATGGCAACACACACACACACACACACACACACACACACA		CATOTOTOTOTAC	H843485	0	4	=	7	3	T19569	609F Homo sapiens cDNA clone 609 similar to SET protein
CATGGCAGACTTCCTCGA H358581 0 0 111 0 0 AA207189 CATGGCAATTCCTCGA H540023 0 3 111 3 1 N80776 CATGGCACTCCCAACT H550274 0 1 11 6 0 AA279492 CATGGCACTCGCAACT H550274 0 1 11 6 0 AA279492 CATGGCACACACACA H656453 0 1 11 0 AA098867 CATGGCACACACACACACACACACACACACACACACACAC	000	CATCINICITY	H114144	0	0	=	⊢	1-	Z36249	HHEA18W H. sapiens partial cDNA sequence; clone HEA18W;
CATGCAGACTCCTGGA H540023 0 3 11 3 1 N80776 CATGGAATTCCTCGA H540023 0 3 11 3 1 N8027809 CATGGACGCCCAACT H550274 0 11 6 0 AA279492 CATGGCGACTGGGG H631275 0 11 1 0 AA8867 CATGGCGACACACACA H656453 0 1 1 0 AA173819 CATGGCACACACACACA H656453 0 1 1 0 A418460 CATGTTGCGGACACTTGA H1022502 0 2 11 2 R48460 CATGTTGCGGACACTTGA H598335 0 7 10 4 9 H41078 CATGCTGCACACTTGAAA H294401 0 1 10 4 9 H41078 CATGCTGCCACTGGGC H1007018 0 1 10 4 9 H41078 CATGCTGCCGAGCA H753656 0 1 10 2	5	CATGCCTGAGTCAG	H358581	0	0	=		<u> </u>	AA207189	2q73e07.r1 Stratagene neuroepithelium (#937231)Homo sapiens cDNA clone 647268 5' similar to TR:E16910 E16910 ENDONUCLEASE.;
CATGGGACCCGAACT HSS0274 0 1 11 6 0 AA025809 CATGGCACCCGAACT HSS0274 0 1 11 6 0 AA028867 CATGGCACCCCCACAG H631275 0 0 11 1 0 AA028867 CATGGCAACCACAG H656453 0 1 11 0 2 R48460 CATGGCAACCACAG H656453 0 1 11 0 2 R48460 CATGGCAACCACAG H656453 0 1 11 0 2 R48460 CATGGCAACACTGAAAA H294401 0 1 10 5 0 H04630 CATGGCACATTGAAAA H294401 0 1 10 5 0 H04630 CATGGCACATTGAAAA H294401 0 1 10 5 0 H04630 CATGGCACACTGCAGG H1007018 0 1 10 5 0 H04630 CATGGCACACACACACA H30618 0 6 1 0 1 1 10 T86566 CATGGTCACACACACACACA H30619 0 6 1 0 1 10 T86566 CATGGTCACACACACACACACA H30619 0 6 1 0 1 10 T86569 CATGGTCACACACACACACACACACACACACACACACACA	3 3	CATGGAATTCCTCGA	H540023	0	~	=	6	-	97.LO8N	za98h04.s1 Homo sapiens cDNA clone 300631 3'.
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CATGGACGCCAACT HSS0274 0 I II 6 0 CATGGACGCCCAACT HSS0274 0 I II 6 0 CATGGCAACACACACACACACACACACACACACACACACA								- '	AA025809	
CATGGACGCCGAACT H550274 0 I II 6 0 CATGGCACCCCAACT H550274 0 I II 6 0 CATGGCACCCCACAG H656453 0 I II 0 2 R48460 CATGGCACCACACAG H656453 0 I II 0 2 R48460 CATGGCACACACACAG H656453 0 I II 2 I L19183 CATGTGCACACATTGA H598335 0 7 I0 4 9 H40730 CATGGCACATTGAAAA H294301 0 I I0 5 0 H04630 CATGGCTGCACACACAG H1007018 0 I I0 4 12 R32331 CATGGTCCCGAGACA H50619 0 6 I 0 2 U03911 CATGTTCCTCGGCA H50619 0 6 I 0 0 2 U03911 CATGGTGAAAAAAA H598380 0 5 I 0 0 2 U03911 CATGGTGAAAAAAA H598380 0 5 I 0 0 1 D55671 CATGGAAATAGGTTTT H12992 0 I I0 6 3 D53402 CATGGAAATAGGTTTT H12992 0 I I0 6 3 D53402 CATGCTGGACCTACC H545906 0 I I0 6 3 D53402 CATGCAAATAGGTTTT H12992 0 I I0 6 3 D53402 CATGCTGGACCTACC H545906 0 I I0 6 3 D53402 CATGCTGGACCTACC H545906 0 I I0 6 3 D53402 CATGCAAATAGGTTTT H12992 0 I I0 6 3 D53402 CATGCACGGCTGGT H371131 0 0 I I 2 T15761					Γ		-	-		2585h05.s1 Soares NbHTGBC Homo sapiens cDNA clone 704313
CATGGACGCCGAACT H550274 0 I II 6 0 CATGGCCGCACCGGG H631275 0 0 II I 0 AA098867 CATGGCGAACACACGG H656453 0 I II 0 2 R48460 CATGGCAACACACGG H656453 0 I II 0 2 R48460 CATGGCAACACACGG H1022502 0 2 II 2 I L19183 CATGGCAGACATTGA H598335 0 7 I0 4 9 H41078 CATGGCAGACATTGA H7598335 0 1 I0 5 0 H04630 CATGGCAGACATTGA H75965 0 2 IO 3 7 S77357 CATGGCAGAAAAAA H753665 0 2 IO 3 7 S77357 CATGGTGCAGAAAAAAA H753665 0 2 IO 3 7 S77357 CATGGTGAAAAAAAA H753665 0 2 IO 3 7 S77357 CATGGTGAAAAAAAA H753665 0 1 IO 0 2 U03911 CATGGTGAGACACACCACACACACACACACACACACACAC									AA279492	31
CATGGGGACCAGGG H631275 0 0 11 1 0 AA098867 CATGGCGACCACAG H656453 0 1 11 0 2 R48460 CATGGGAACACACAG H656453 0 1 11 0 2 R48460 CATGGTGGAACACACAG H1022502 0 2 11 2 1 L19183 CATGTTGCGGACATTGA H598335 0 7 10 4 9 H41078 CATGGCAGACATTGA H794401 0 1 10 5 0 H04630 CATGGCAGACATTGA H794401 0 1 10 5 0 H04630 CATGGCAGACATTGA H79565 0 1 10 4 12 R2331 CATGGTGAAAAAAA H753665 0 2 10 3 7 S77357 CATGGTGAAAAAAAA H753665 0 2 10 3 7 S77357 CATGGTGACACACACA H506149 0 6 10 6 1 M34338 CATGGTGACACACACACA H506149 0 6 10 0 2 U03911 CATGGTGAAAAAAAA H12992 0 1 10 6 3 D53402 CATGGACCACTACC H545906 0 1 10 6 3 D53402 CATGGACCACTACC H545906 0 1 10 6 3 D53402 CATGGAAATAGGTTTT H12992 0 1 10 6 3 D53402 CATGGAAATAGGTTTT H12992 0 1 10 6 3 D53402 CATGGAAATAGGTTTT H12992 0 1 10 6 3 D53402 CATGCACGCGCGCGCGCGCGCGCGCGTGGT H371131 0 0 10 1 2 T35761	_	CATGGACGCGAACT	H550274	0	F	=	9	0		Unknown
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CATGGGAACACACAG H656453 0 1 11 0 2 R48460 CATGGGAACACACACAG H1022502 0 2 11 2 1 L19183 CATGGTGAGCACACTGAAA H1022502 0 2 11 2 1 L19183 CATGGCAGACATTGA H598335 0 7 10 4 9 H41078 CATGGCAGACATTGA H19435 0 0 10 24 0 R77027 CATGGCTTGGCAGG H1007018 0 1 10 4 12 R32331 CATGGTTGCAGGC H1007018 0 1 10 4 12 R32331 CATGGTTGCTGGAGC H1007018 0 1 10 4 12 R32331 CATGGTTGCAGGC H1007018 0 1 10 4 12 R32331 CATGGTTGCAGGC H1007018 0 1 10 4 12 R32331 CATGTTCTCTGGGC H1007018 0 1 10 4 12 R32331 CATGGTTGCAGGC H1007018 0 1 10 4 12 R32331 CATGGTTGCTGCAGG H1007018 0 1 10 4 12 R32331 CATGGTTGCTGCAGC H150619 0 6 10 6 1 M34338 CATGGTGAAAAAAA H753665 0 1 10 0 2 U03911 CATGGTGAAATAGGTTTT H12992 0 1 10 6 3 D53402 CATGGAAATAGGTTTT H12992 0 1 10 2 10 3 T51516		LATGGGGGGCTGGGG	H631275	0	0	=	_		AA098867	489535 3' similar to SW:A5_XENLA P28824 A5 PROTEIN PRECURSOR
H1022502 0 2 11 2 1 L19183 H1022502 0 2 11 2 1 L19183 H598335 0 7 10 4 9 H41078 H294401 0 1 10 5 0 H04630 H719435 0 0 10 24 0 R77027 H719435 0 0 10 24 0 R77027 H719435 0 1 10 4 12 R32331 -497192 0 8 10 1 10 T86566 H753665 0 2 10 3 7 S77357 H506149 0 6 10 6 1 M34338 H506149 0 6 10 6 1 M34338 H242380 0 5 10 9 7 D55671 H242380 0 5 10 9 7 D55671 H345906 0 1 10 3 1 J03569 H345906 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D53402 H37131 0 0 10 1 2 T35761	_	CATGGGAACACAG	H656453	0	-	=	0	2	R48460	yj67c12.r1 Homo sapiens cDNA clone 153814 5'.
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H598335 0 7 10 4 9 H41078 H294401 0 1 10 5 0 H04630 H719435 0 0 10 24 0 R77027 H1007018 0 1 10 4 12 R32331 -497192 0 8 10 1 10 T86566 H736649 0 6 10 6 1 M34338 -835515 0 1 10 0 2 U03911 H242380 0 5 10 9 7 D55671 H345906 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D53402 H37131 0 0 10 1 2 T51971		ראומווחרמטימייי				T	-		H61710	yr24a07.s1 Homo sapiens cDNA clone 206196 3'.
H598335 0 7 10 4 9 H41078 H294401 0 1 10 5 0 H04630 H719435 0 0 10 24 0 R77027 H1007018 0 1 10 4 12 R32331 -497192 0 8 10 1 10 T86566 H753665 0 2 10 3 7 S77357 H756149 0 6 10 6 1 M34338 -835515 0 1 10 0 2 U03911 H242380 0 5 10 9 7 D55671 H545906 0 1 10 3 1 J03569 H12992 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D51402 H371131 0 0 10 1 2 T35761						T	t	-	H77330	yul If12.s1 Homo sapiens cDNA clone 233519 3'.
H598335 0 7 10 4 9 H41078 H719435 0 1 10 5 0 H04630 H719435 0 0 10 24 0 R77027 H1007018 0 1 10 4 12 R32331 -497192 0 8 10 1 10 T86566 H753665 0 2 10 3 7 S77357 H76149 0 6 10 6 1 M34338 -835515 0 1 10 0 2 U03911 H242380 0 5 10 9 7 D55671 H345906 0 1 10 3 1 J03569 H12992 0 1 10 6 3 D53402 H12992 0 1 0 6 1 D61243						T	\vdash	\vdash	N69482	za18d05.s1 Homo sapiens cDNA clone 292905 3'.
H1294401 0 1 10 5 0 H04630 H719435 0 0 10 24 0 R77027 H1007018 0 1 10 4 12 R32331 -497192 0 8 10 1 10 T86566 H753665 0 2 10 3 7 S77357 H753645 0 6 10 6 1 M34338 H26149 0 6 10 6 1 M34338 H242380 0 5 10 9 7 D55671 H242380 0 1 10 0 2 U03911 H345906 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D53402 H12992 0 1 0 6 1 D61243	7771	CATCCCAGACATTGA	H598335	0	1	2	4	6	H41078	yp52c11.s1 Homo sapiens cDNA clone 191060 3' simil
H119935 0 0 10 24 0 R77027 H1007018 0 1 10 4 12 R32331 -497192 0 8 10 1 10 T86566 H753665 0 2 10 3 7 S77357 H506149 0 6 10 6 1 M34338 -835515 0 1 10 0 2 U03911 H242380 0 5 10 9 7 D55671 H37596 0 1 10 5 3 D53402 H12992 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D53402 H371131 0 0 10 1 2 T51971	3 5	CATGCACTTGAAAA	H294401	0	-	2	2	0	H04630	yj49g03.r1 Homo sapiens cDNA clone 152116 5'.
H1007018 0 1 10 4 12 R32331 -497192 0 8 10 1 10 T86566 H753665 0 2 10 3 7 S77357 H506149 0 6 10 6 1 M34338 -835515 0 1 10 0 2 U03911 H242380 0 5 10 9 7 D55671 H545906 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D53402 H371131 0 0 10 1 2 T51971		CATGGGTTGGCAGG	H719435	0	0	2	24	0	R77027	yi66e12.r1 Homo sapiens cDNA clone 144238 5'.
-497192 0 8 10 1 10 T86566 H753665 0 2 10 3 7 S77357 H566149 0 6 10 6 1 M34338 -835515 0 1 10 0 2 U03911 H242380 0 5 10 9 7 D55671 H545906 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D53402 H12992 0 1 10 6 3 D61243 H371131 0 0 10 1 2 T35761		CATGTTCCTCGGGC	H1007018	0	-	2	-	12	R32331	yh68g02.s1 Homo sapiens cDNA clone 134930 3' simil
H753665 0 2 10 3 7 S77357 H506149 0 6 10 6 1 M34338 -835515 0 1 10 0 2 U03911 H242380 0 5 10 9 7 D55671 H345906 0 1 10 3 1 J03569 H12992 0 1 10 6 3 D53402 H12992 0 1 0 6 3 D53402 H37131 0 0 10 1 2 T5191		CATGCTGCGAGCT	-497192	0	∞	01	_	9	T86566	yd77g07.r1 Homo sapiens cDNA clone 114300 5' simil
H506149 0 6 10 6 1 M34338 -835515 0 1 10 0 2 U03911 H242380 0 5 10 9 7 D55671 H545906 0 1 10 3 1 J03569 H12992 0 1 10 6 3 D53402 H12992 0 1 0 6 3 D53402 H37131 0 0 10 1 2 T51761		CATGGTGAAAAAA	H753665	0	2	01	3	7	S77357	transcript ch 1 1 [human, RF1, RF48 stomach cancer c
.835515 0 1 0 2 U03911 H242380 0 5 10 9 7 D55671 H545906 0 1 10 3 1 J03569 H12992 0 1 10 6 3 D53402 H240131 0 0 10 1 16 1 H37131 0 0 10 1 2 T35761	_	CATGCTGTGCAGCA	H506149	0	9	0	9	_	M34338	Human spermidine synthase
H124380 0 5 10 9 7 D55671 H545906 0 1 10 3 1 J03569 H12992 0 1 10 6 3 D53402 T61971 H371131 0 0 10 1 2 T35761		CATGTAGTTTGTGG	-835515	0	_	01	0	7	U03911	Human mutator gene (hMSH2)
H12992 0 1 10 3 1 J03569 H12992 0 1 10 6 3 D53402 T61971 H371131 0 0 10 1 2 T35761		CATGATGTAGTAGTG	H242380	0	2	으	6	7	DSS671	Human heterogeneous nuclear ribonucleoprotein
H12992 0 1 10 6 3 D53402 T61971 H371131 0 0 10 1 2 T35761	7	CATGGACCCACTACC	HS45906	0	E	2	-		103569	Human lymphocyte activation antigen 4F2 large subunit
H371131 0 0 10 1 2 T35761	7,	CATGAAATAGGTTIT	H12992	0	E	2	9	3	D53402	Human fetal brain cDNA 5'-end GEN-108D03.
H371131 0 0 10 1 2 T35761									T61971	yb96f02.rl Homo sapiens cDNA clone 79035 5'.
H371131 0 0 10 1 2 T35761								-	D61243	Human fetal brain cDNA 5'-end GEN-171G06.
H371131 0 0 10 1 2 T35761									N77240	yv44d02.r1 Homo sapiens cDNA clone 245571 5'.
	1.7	177 CATGCCGGGCGTGGT	H371131	0	0	9	片	7	T35761	EST90898 Homo sapiens cDNA 5' end similar to EST c

Treated to many capiene CONA S' end similar to None.	H555168 0 8 10 3 3 131901 ES140/19 HONING SEPTING SECTION SECT		15234	x08764 [IHSMPP4] H.sapiens mRNA for M-phase phosphoprotein, inpp4, 13230p	LOTO/V	2000	Oliklicani	The Day of the Day of the Angel of the Day o	U8/433 Human minist for Night to Bene, Permis		
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	H\$55168				H648]		123007	11272021	2 0 0 0 0 0 0	H0110H	
	CTTO A CATO A COST A CO	CALCOACTOACTTO			TAACCCCAAACTCCCCC	- パー・パー・パー・パー・パー・パー・パー・パー・パー・パー・パー・パー・パー・パ		- AN INTERIOR CONTROL OF THE		Les La Tanana CATACA TANA	101 (00,000,000,000,000,000,000,000,000,000

Table ? - Transcripts decreased in colon cancer

Transcripts decreased in only colon primary tumors

compared to normal colon (51 genes)

NC: Normal Colon

TU: Colon Primary Tumor CL: Colon Cancer Cell Line PT: Pancreatic Primary Tumor PC: Pancreatic Cancer Cell Line

	_	1	η-	_	_	1		-,			_	_	т	Т	\neg	\neg	1	_	Т	Т	Т	Т	Т		_		_	\neg
Gene Name	Unan mDMA for heta actin	numaii iii Nava 101 octa-acuii.	Human mKNA for cytosketetal gamina-actini.	Human mRNA for cytokeratin 18.	Human lipocortin II mRNA.	Human mRNA for calcium dependent protease (small subunit)	H.sapiens CpG island DNA genomic Msel fragment, cl	zd30d02.r1 Soares fetal heart NbHH19W Homo sapiens	Human fetal brain cDNA 5'-end GEN-141D02.	Unknown	Human thyroid hormone binding protein (p55) mRNA,	w05d05 s1 Homo saniens cDNA clone 270345 3'	1) Control of Control of Long Content	2006aUS.rl Soares retai lung indiricity w moine sapiens	Human mRNA for argininosuccinate synthetase.	Human mRNA for very-long-chain acyl-CoA dehydrogen	Human keratinocyte cDNA, clone 173.	himan alnha-hihilin mRNA. 3' end.	Manifest and Total Cold Library II Home canions CDNA Cond	AA341033 ES14/106 Fedal Kidney II Holing Saprens Color S and	H.sapiens Id! mKNA.	H.sapiens mRNA for BiP protein.	Human cytochrome c oxidase subunit VIII (COX8) mRNa	Human Na, K-ATPase alpha-1 subunit mRNA, complete c	OBINE SUSSUIR SOLD SISSUED SOLD SOLD SOLD SOLD SOLD SOLD SOLD SOL	Edicopolitication provider CDMA clone 153030 5'	yjoycou. I nomo sapiens con a cione i coca	Human Heart cDNA, clone 3NHC0642.
Accession	_13	700351	X04098	502 X12883	D00017	X04106	265513	W61077	D60944		102783	N33047	TOCOL	W07627	X01630	D43682	D29146	75007	VCCOON .	AA341633	X77956	X87949	104823	U16798	050350	25057	KSUU13	C02981
۲	:	=	2	202	104	4	32	32	4	~	, 2	: 5	3	8	2	∞	~	۶	٤	≖	이	2	=	₽	5	3		
5		2	8	36	S	37	0	٥	~	2	3 5	1	2	24	2	=	12	: -	٠	2	8	13	9	0	٤	2		
5	3	2	8	245	38	88	\$	2	1 %	3 7	; ;	1	*7	76	57	92	ير	3 8	3	<u>∽</u>	36	14	2	~	: ;	Ş		
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	~+	84	170	137	25	15	: =	: 5	3 5	; ;	;	ş	\$	43	42	8	Ę) 	× ×	37	35	33	33	E	3 3	22		
H	٥	H654591	H468434	H263478	H413181	11278022	11591074	11504008	1177040	040/74H	H349801	H38/10/	H621140	H150053	H28235	H615802	13903011	1200001	H648575	H955615	H456167	H937452	H755160	11026031	1 C0070H	H760267		
	# Tag sequence	1 CATGGCTTTATTTGT	2 CATGCTAGCCTCACG	2 CATCOLLA A A COLATIONA	S CAIGCAAACCAICCA	4 CAIGCIICCAGCIAA	S CATGCCCCAGIIOCI	6 CATGGAIGACCCC	7 CATGCIGIACAGACA	8 CATGCGGACTCACIG	9 CATGCCCCCGCGGAA	10 CATGCCTGGAAGAGG	11 CATGGCCTGGCCATC	12 CATGAGGAGGAGGAG	12 CATOLANCETOCAGGG	LA CATGRACOTOCAGO	14 CATOUCCUCCIOCA	15 CATGTGGGGAGAGGA	16 CATGGCTGCCCTTGA	17 CATGTGGCCATCTGC	19 04 TOCOTTOCOG	S CATOCOTOTO S	19 CATGLOCALCIOGIC	20 CATGGTGACCTCCTT	21 CATGTAGCTCTATGG	22 CATGGTGCGCTAGGG		

								loning to similar to ubiquinol
			-					EST30445 Homo sapiens CDINA 3 end similar to conquired
UULULUUUUUUU T	H694767	28	9	20	9	\neg	T31329 C	Cytochrome-c reductase, v.4 A.D.a.
23 CATGGGGGGGGTGGG	H382130	27	9	12	3	61	ונ	Unknown
24 CATGCCICCAGIAC	76728617	12	-	4	000	7 H	H63643 y	yr34d11.r1 Homo sapiens cDNA clone 20/189 5 simil
25 CATGCCTGIGACAGE	1700001	2		∞	12	<u>></u>	W60924 z	zd27c08.rl Soares fetal heart NbHH19W Homo sapiens
26 CATGTCACAGIGCUI	1000000	1 5	-	-	=	13	L25081	Human GTPase (rhoC) mRNA, complete cds.
27 CATGAATAAAGGCIA	020160111	3 5	-	=	5	25 D	D45887	Human mRNA for calmodulin, complete cds.
28 CATGTTGTTGAA	H1031929	3 2	, -	<u> </u> =	╀	1	Γ	yy66b11.s1 Homo sapiens cDNA clone 278493 3.
29 CATGAAGGTAGCAGA	11740117	3 7	- ~	: 0	╀╌	1	R68653	yi14b06.s1 Homo sapiens cDNA clone 139187 3.
30 CATGGTGTTGGGGG	1102021	; -	1-	-	-	5 ×	X90858	H.sapiens mRNA for uridine phosphorylase.
31 CATGTGCAGCGCCTG	H930344	7 6	- -	, -	-	1		yn54c02.s1 Homo sapiens cDNA clone 172226 3' simil
32 CATGATGGCACGGAG	H238697	3 6	, -	- 1	, -	1		EST17149 Homo sapiens cDNA 5' end similar to None.
33 CATGGCCAGACACCC	H008320	3 8	- -	, [-	-	-	Γ	Human gene for alpha 1 globin.
34 CATGCTTCTTGCCCC	H515990	3	,		, [5	Т		Human jun-B mRNA for JUN-B protein.
15 CATGACCCACGTCAG	H86453	2	7	1	*	7		vignen8.s1 Homo sapiens cDNA clone 156038 3.
14 LATGRETICETICE	H686458	<u>∞</u>	_	4	7	Т	T	STATION Home saniens cDNA clone 153787 3.
2000				7	1	<u> </u>		Sactor of them caniene o'DNA clone 154253 3.
				-		×.		1/2003.51 nolino sapiens conversion of clarks III is
CHOCOCOC	USKIKKO	~	7	7	9	X 91	X12910	a+,K+ Al Pase gene exons 1 - 3 (alpula 111
37 CATGGAGGGCCGGIG	200/061	: :	-	-	,	-		Unknown
38 CATGGATGAATCCGG	H581847		-\	\ \ -	1	Т	X81006	H. sapiens HCG I mRNA.
39 CATGAGCCCGACCAC	H153109	اء	1	= :	-	Τ.	Γ	Homo sapiens porin (por) mRNA, complete cds and tr
10 CATGGTTCAGCTGTC	H774780	9]	7	١-	1	┰	T	Human 78 kDa gastrin-binding protein mRNA, complet
1 CATGCTCGCTCAGT	H383443	16	-	~		T	T	11 DENE DONA partial cde
41 CATOCOTOCOTOCOTOCOTOCOTOCOTOCOTOCOTOCOTO	H265219	15	_		٥	히		Human Deive Hillary, paintal cas.
42 CATOCARATARKICO	H940378	2	-	∞	0	~		Human semaphorin v innaka, compress cas.
43 CA1610CCCCCCA	11601752	2	0	9	4	3	D12038	Human HepG2 3'-directed Mbol CDINA, Clolic 31.30.
44 CATGCCAGTGGCCIC	10011		-	-	~	<u>~</u>	U77396	Human TNF-alpha inducible responsive element mKNA,
45 CATGCTGGCCTGAA	151705H	: :	-	1	=	12	Z29093	H.sapiens EDDR1 gene for receptor tyrosine kinase.
46 CATGGCCCATTGGAG	H611303	2 5	- -	,	:	7		ye38a04.s1 Homo sapiens cDNA clone 119982 3'.
47 CATGAAGAAACCTC	H32792	2	>	,	1	Т		za25g05.s1 Homo sapiens cDNA clone 293624 3'.
					T			zb86e03.s1 Soares senescent fibroblasts NbHSF Homo sapiens cDNA
							N98502	clone 310492 3'
		!	٩	1	1	1	F18838	H.sapiens EST sequence (007-X1-01) from skeletal m
48 CATGGAATGATTTCT	H538878	2	>	-	, 	1		zr21b10.s1 Stratagene NT2 neuronal precursor 937230 Homo sapiens
	116.21.23.	2	_	~	~		AA226928	cDNA clone 664027 3'
49 CATGGCCTGGTCCTI	H021219	1	0	-	-	Γ	M60047	Human heparin binding protein (HBp17) mRNA
SO CCATGGCCCACACAG	22201							

zc45e09.rl Soares senescent fibroblasts NbHSF Homo SI CATGGGATTCCAGTT

Transcripts decreased in both colon primary tumors and colon cancer cell lines compared to normal colon (130 genes)

NC: Normal Colon TU: Colon Primary Tumor

CL. Colon Cancer Cell Line PT. Pancreatic Primary Tumor PC. Pancreatic Cancer Cell Line

			2	E	=	7	24	Accession	Gene Name
=	Tag Sequence	Tag Number	١		_	-		Caption	Il. man mDNA for eviolentin 8
1-	CATCCTCAGCTAC	H382109	803	<u>5</u>	304	36	ĝ	X12882	חוווווווו ווווווווו ליינייוו כי
-]	CATOCTAGACTTCA	H460926	80	282	402	142	497	F15636	H.sapiens mitochondrial ES1 sequence (002113)
1	CALGOLOGICACO	H610997	705	88	7	7	_		Unknown
	CATGGCCCAGGTCAC	100002	\$12	348	2	5	235	F16940	H.sapiens mitochondrial EST sequence (U09-11-21)
7	1 CATGACCCI IGGCCA	1191562	Ş	S	4	0	6	M10050	M10050 Human liver fatty acid binding protein (FABP) mRNA
ς.	SCATGACATTGGGIGA	00/00/		: 2	1	S	=	\$61953	c-erbB3=receptor tyrosine kinase (alternatively sp
9	6 CATGGCGAAACCCTG	H62268U	260	3 5	15	╅	į		H.sapiens mitochondrial EST sequence (1-t-02) from
7	CATGAGCCCTACAAA	H133301	2 5		٥	╈	6		va04c01.r2 Homo sapiens cDNA clone 60480 5'.
œ	8 CATGGACCCAAGATA	H242828	\$		<u>,</u>	1	T		v/4/1a01.s1 Homo sapiens cDNA clone 160776 3'.
				1	1	1	T	_	HUMGS02706 Human colon 3'directed Mbol cDNA, HUMGS02706,
					-			025586	725586 clone cm 1673.
				1	1	1	T	106160	Trociso Manager Homo caniens CONA clone 117195 3.
L					1	1		190100)(U)(UE:31 10110 3-1111 1011
١	O TOUCHOUSE	H617195	256	8	148	144	28	X64364	X64364 H.sapiens mkna 10r Mo annigen.
`	CALGOCCOCC	H1076814	202	75	84	235	369	M11146	M11146 Human ferritin H chain mKNA, complete cus.
2	10 CA101100001115C	LA70577	ž	2	0	=	~	L15203	L15203 Human secretory protein (P1.B) mRNA, complete cds.
Ξ	11 CATGCTCCACCCGAA (of U)	11000	3	2	-	2	2	X93036	X93036 H.sapiens mRNA for MAT8 protein.
12	12 CATGGCAGGGCCTCA	H6000/0	2	8	,	:	:		vv07h09.r1 Homo sapiens cDNA clone 242081 5' similar to SP:A39484
		11274023	104	24	66	40	39	H93844	A39484 ANDROGEN-WITHDRAWAL APOPTOSIS PROTEIN RVPI,
=1	13 CATGATCGTOGCGGG	11271574	8	g	[⊡	m	8	F17001	H.sapiens mitochondrial EST sequence (011-T1-13) f
7	14 CATGCAAGCAICCC	101/21		1	×	5	20	V00503	V00503 Human mRNA for keratin 19.
15	15 CATGGACATCAAGTC	H544012	è	3	2	1			2605a11.rl Soares fetal lung NbHL19W Homo sapiens cDNA clone
<u> </u>									301148 5' similar to gb: V00567 BETA-2-MICROGLOBULIN
	* * <u>1.1.</u>	H782013	178	110	4	340	139		W16632 PRECURSOR (HUMAN);.
2	16 CATGGTTGTTAA								zo31h04.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
								AA143804 588535 3	588535 3'

					-			07 7102h07 s1 Stratagene colon (#937204) Homo sapiens cDNA clone
				_			AA133597 512115 3	5121153
		-	T		1		T53199	T53199 ya86c05.s1 Homo sapiens cDNA clone 68552 3'.
11 CTAGTGCTCCTACCC	H947654	174	77	-	0	0	R00081	ye73c04.s1 Homo sapiens cDNA clone 123366 3'.
10 CATCACCCTGATG	H284132	172	2	92	~	9	M16364	M16364 Human creatine kinase-B mRNA, complete cds.
18 CALOCACCACACACACACACACACACACACACACACACAC			Γ					yf22e12.s1 Homo sapiens cDNA clone 127630 3' similar to contains Alu
	H368200	163	9	4	2	4	R09410	repetitive element
200000000000000000000000000000000000000			Γ					HUMGS0003915, Human Gene Signature, 3'-directed cDNA
							C01918	
				T		\vdash		yq04h09.s1 Homo sapiens cDNA clone 196001 3' similar to
							R92735	contains Alu repetitive element
								zh78e12.s1 Soares fetal liver spleen INFLS S1 Homo sapiens
								cDNA clone 418222 3' similar to contains Alu repetitive element
2 CATOCOCTOC	H501111	163	2	0	92	_	j	H.sapiens pS2 protein gene.
20 CATOCCCCTCGATC	H350116	09	8	24	88	181	M18981	Human prolactin receptor-associated protein (PRA)
21 CATOCCCCTOON C	H1001401	091	74	2	74	71	M64303	Human galactoside-binding protein mRNA.
22 CAIGITCACIOLORS	H256186	155	2	-	=	9	X16455	Human mRNA for carcinoembryonic antigen pCEA80-11.
23 CATUAL IGGAUINCI	H403039	149	4	32	8	37	U14943	Human MHC antigen (HLA-B) mRNA, complete cds.
24 CAIGCIGACCIGIOI	H149715	145	S	8	 —	23	M81457	M81457 Human calpactin 1 light chain mRNA, complete cds.
25 CATGAGCAGATCAGG	LLPSSYH	12	۳	-	75	9	C21047	HUMGS0002546, Human Gene Signature, 3'-directed cDNA sequence
26 CATGGGAAAACAGAA	10000			T		+-		2021h08.s1 Stratagene colon (#937204) Homo sapiens cDNA
							4A132779	AA132779 clone 587583 3' similar to SW. LEG4_RAT P38552 GALECTIN-4
				T				zl68h06.s1 Stratagene colon (#937204) Homo sapiens cDNA
							4A054072	AA054072 clone 509819 3'
								zo18g08.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
		_			_		AA132736	AA132736 587294 3' similar to SW:LEG4 RAT P38552 GALECTIN-4
22 CATOTOACTOAG	H857781	122	2	7	2	7	X04412	Human mRNA for plasma gelsolin.
21 CATOTOCACOATOCAG	H936217	122	78	32	25	2	X77658	H. sapiens mRNA for HLA-B*7301.
28 CATOLOCACO								zo35c09.si Stratagene colon (#937204) Homo sapiens cDNA clone
O TO TO TO TO TO A VITILITY OF A	H657337	115	7	-	4	7	AA146606 S88880 3	588880 3'
62								zo35g09.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
-							AA146775 588928 3	588928 3
								zo74g11.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
							AA161043 592676 3'	592676 3'
						ŀ		

Į				+	\vdash	-	\vdash		z183108.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
							₹	AA088704 511239 3'	511239 3'
Ş	CATCCGAGGGGCCAG	H404117	=	32	54	9	\$	H00427	yj23g11.r1 Homo sapiens cDNA clone 149636 5'.
				-		-	-		2063d03.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
				1	+	\dashv	<u> </u>	$\overline{}$	091007 3
			_		_		_	T08562	EST06454 Homo sapiens CUNA Clone HIBBUST 3 CITU.
									zm21a12.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
		7				-	₹	S	526270 3'
=	CATGTAAATTGCAAA	H790417	113	9	_	0		X73502	H. Sapiens mRNA for cytokeratin 20.
Τ.	CATCCCTCCCCC	H686762	-13	36	48	45 '	43	103191	Human profilin mRNA, complete cds.
7 7	CATGGTGCTGAATGG	H761359	80	20	30	1 19]	U02629	Human smooth muscle myosin alkali light chain mRNA
	24 CATGOTGCACTGAGG	H758243	5	2	36	34	82	X07059	Human M4-50 mRNA for HLA class I antigen.
,	CATOOL ACCOUNT	H1032614	107	=	4	3	37	F15592	H.sapiens mitochondrial EST sequence (001 T24) from
	2222224		T	T	\vdash		-		zl74e07.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
		H157779	106		-	۳	<u>¥</u>	A053660	AA053660 510372 3' similar to contains Alu repetitive element
٥	CAIGCCCICCGAAG	771,75011		+	+	+	T		HUMGS04077 Human colon 3'directed Mbol cDNA, HUMGS04077,
							-	D25711	clone cm 1210
1			1			+	\vdash		H.sapiens CpG DNA, clone 140c4, reverse read cpg 14(Mitochondria
	**************************************	H178755	105	-2	77	4	27 /	256800	EST
7 2	3) CATCATACTCACTC	H204104	102	=	0	0	0	M95174	Human guanylin mRNA, complete cds.
3	CATOAICCOCTOGG	H484987	≘	25	~	4	91		Unknown
25	יא ראומרובמרמבומממ			T		\vdash	\vdash		yn01b01.r1 Homo sapiens cDNA clone 167113 5' similar to SP.ZK783.1
-		H697514	82	32	78	37	- 65	R90863	CE00760 ;.
3	222222222222222222222222222222222222222				\vdash			T24702	T24702 EST277 Homo sapiens cDNA clone 10H4.
]=		HS33666	8	55	42	78	. 28	X95404	X95404 H.sapiens mRNA for non-muscle type cofilin.
÷ (5	4) CATGCCAGGGGAGA	H338569	25	22	78	30	91	X67325	X67325 H.sapiens p27 mRNA.
; 5	CATGACAGGCAAGA	H70211	74	31	30	01	31	F16604	H.sapiens mitochondrial EST sequence (009128) from
									za16a03.s1 Homo sapiens cDNA clone 292684 3' similar to contains Alu
-	14 CATCAGA ATAGCTTG	H134304	69	53	_	~	0	N69361	
7 7	0.1000.000.000.000			T	T		-		ze30b10.s1 Soares retina N2b4HR Homo sapiens cDNA clone
							<u> </u>	A015918	AA015918 360475 3' similar to contains Alu repetitive element
		·				-			yll4h01.s1 Homo sapiens cDNA clone 158257 3' similar to contains Alu
								H26689	repetitive element; contains TARI repetitive element ;.
_						-			2779h I I.s i Soares NhHMPu S1 Homo sapiens cDNA clone 681957 3
	TOBOUTOLOGOT	H424875	89	6	9	~	23 A	A256365	AA256365 similar to WP:C33A12.7 CE05353
}]	CAIGCOCIOIOGGG				1				

					+	-	<u> </u>	2010e11 el Soares senescent fibroblasts NbHSF Homo sapiens cDNA
						W4	W47357	clone 324716 3'
					T			zb90(03.s1 Soares senescent fibroblasts NbHSF Homo sapiens cDNA
				,		WI		clone 310877 3'
						R0,	R07159	yf13h12.s1 Homo sapiens cDNA clone 126791 3'.
46 CATGCATAGGTTTAG	H314109	88	~	0	0	0 L02	L02785	Homo sapiens colon mucosa-associated (DRA) mRNA
47 CATGGCGACCAGGT	H614731	65	<u>≎</u>	0	3	110 9	U11862	Human clone HP-DAO1 diamine oxidase
48 CATGAGCTCTTGGAG	H161769	8	=	-	-	2 N93	N93240 z	zb68b06.s1 Homo sapiens cDNA clone 308723 3'.
							=	NIB1986 Normalized infant brain, Bento Soares Homo sapiens cDNA
		-				T16	T16906 3	3'end.
							Î	yu22h07.s1 Homo sapiens cDNA clone 234589 3' similar to
						H78	H78256 S	SP:SBP_MOUSE P17563 SELENIUM-BINDING
					-		=	EST47523 Homo sapiens cDNA 3' end similar to similar to Selenium-
						T32	T32362 b	binding protein liver.
49 CATGCCAACGCGCT	H344474	52	-	0	3	0 000	V00493 F	Human messenger RNA for alpha globin.
SO CATGGACGCGCGCG	H550554	S	21	2	7	14		Unknown
SI L'ATGACCCCCCCCCC	H87386	54	91	15	15	3 XSI	X51346 F	Human jun-D mRNA for JUN-D protein.
S CATGATGCGGAGAA	H236169	22	9	2	=	7 R34		yh83f04.r1 Homo sapiens cDNA clone 136351 5'.
75					-	H03	H03961 y	yj44e07.s1 Homo sapiens cDNA clone 151620 3'.
			Γ			R33	R33498 y	yh83104.s1 Homo sapiens cDNA clone 136351 3'.
					-	_		zi71e06.rl Stratagene colon (#937204) Homo sapiens cDNA clone
st CATGTCAGCTGCAAC	11862097	2	9	0	0	0 AA0	53043	AA053043 510082 5'
SA CATGGTAAGTGTACT	H723890	8	14	15	_	30 F17		H.sapiens mitochondrial EST sequence (007T13) from
\$\$ CATGTGTGGGTGCTG	H977640	49	70	17	21	8 213		H.sapiens mRNA for E-cadherin.
\$6 CATGGCTGTGCCTGG	H650847	48	12	15	8	31 X15		Human mRNA for pancreatic trypsinogen III.
57 CATGTGAGTGACAGA	H929299	48	4	0	0	\dashv	H14641	yl26g02.s1 Homo sapiens cDNA clone 159410 3:
\$8 CATGGGCTGGGCCTG	H686744	47	=	=	32	8 M20	9469	M20469 Human brain-type clathrin light-chain b mRNA,
							_	yy92c07.s1 Homo sapiens cDNA clone 281004 3' similar to contains Alu
59 CATGTAATCCCAGCA	H800074	46	~	~	∞	- NS0		repetitive element; contains element MER32 repetitive element
60 CATGGACCAGTGGCT	H545514	45	-	٥	히	-	U79725	Human A33 antigen precursor mRNA, complete cds
61 CATGGGCACCGTGCT	H673210	44	의	-	픠	-		Unknown
62 CATGAAGGACCTTT	H41344	43	2	4	2	24 HII		ym14f06.r1 Homo sapiens cDNA clone 47991 5.
						HS		yt85h08.s1 Homo sapiens cDNA clone 231135 3'.
		\prod	П	\prod	\sqcap	T4(T40539 y	ya05b02.s1 Homo sapiens cDNA clone 60555 3'.

							AA	303091 E	AA303091 EST12940 Uterus tumor 1 Homo sapiens cDNA 3' end
					├-	├		7	2a52d02.rl Soares fetal liver spleen INFLS Homo sapiens cDNA clone
(V)	CATGGCAGCTCCTGT	H599903	43	∞	=	24	<u>×</u>	02429 4	W02429 [296163 5]
							Ż	20325	N20325 yx44ci1.s1 Homo sapiens cDNA clone 264396 3.
1						-	ž	N45127 y	yz13c12.s1 Homo sapiens cDNA clone 282934 3'.
1			T			\vdash	_	-	zb38c11.s1 Soares parathyroid tumor NbHPA Homo sapiens cDNA
							ž	N90407 c	clone 305876 3'.
_	CATCOTOTOTO	H972720	5	12	=	22	S O	U03106	Human wild-type p53 activated fragment-1 (WAF1) mR
5	21.00.00.00				\vdash	-	-		zc1101.s1 Soares parathyroid tumor NbHPA Homo sapiens cDNA
	A J J J J J J A A A J A D T A D	H65878	42	91	7	- 2	11 W.	W37827 c	clone 322009 3*
ر اد	CONCARD CONCARD IN				\vdash		-	30	gblW15332 W15332 zc16d10.s1 Soares parathyroid tumor Nb11PA
_							≥	W15332	Homo sapiens cDNA clone 322483 3'
				1		\vdash	-	1	zc04g10.s1 Soares parathyroid tumor NbHPA Homo sapiens cDNA
							<u>≯</u>	W32410	clone 321378 3'
				T	\dagger	t	Z	N32312 V	vw82c01.s1 Homo sapiens cDNA clone 258720 3'.
		1878111	14	9	=	9	0	T	Human sodium/potassium-transporting ATPase beta-3
ز 99	66 CATGIAGGATGGGG	0133011			-	╀	<u>ا</u> م	Т	Unknown
67 CA	CATGACTGTGGCGGC	H126619	7	1	+	+	+	1	14611 -1 Charles amicals 037300 Homo caniene CONA clone
				,	:			7 2 0 0 0 1	2p4411.31 Stratagene muscle 93/209 moins saptens control and 1908 (413333 3) similar to contains Alu renetitive element:
V) 89	CATGGTAGCAGGTGT	H730287	₽	1	2	╌	<u>د</u>	210001	187-00 et Homo ganiens CDNA clone 136734 3' similar to contains Alu
		-							yllo/cod, st. 110110 sapicits colors design and a sapicity of the sapicity of
						1	¥	K34090	repetitive eletitetiti.
1						_			yh87e04.s1 Homo sapiens cDNA clone 136734 3' similar to contains Alu
						_	~	R34696 I	repetitive element;.
									zq06e03.s1 Stratagene muscle 937209 Homo sapiens cDNA clone
							AA	194497	AA194497 628924 3' similar to contains Alu repetitive element
						-	_		hbc760 Homo sapiens cDNA clone hbc760 3'end similar to nonspacific
- 9	**************************************	H53508	40	12	0	<u>-</u>	1 O	T11144	crossreacting antigen.
<u>ဒ</u> ါ	CALGAAICACAAAIA	2000011					-		zl67e01.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
							AA	058357	AA058357 509688 3' similar to TR:G189087
)	C05803	similar to none
+						T		_	zo31e02.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
		H167606	40	=	4	4	S AA	143765	AA143765 588506 3'
) -	CATGAGGATGGTCCC				1				zp45b09.s1 Stratagene HeLa cell s3 937216 Homo sapiens cDNA clone
							¥ ¥	179299	AA179299 612377 3'
					1				

	M35252 Human CO-029.	R87448 ym89c10.s1 Homo sapiens cDNA clone 166098 3'.	X79882 H.sapiens Irp mRNA.	_	J03037 Human carbonic anhydrase II mRNA, complete cds.	_1	M92843 [H.sapiens zinc finger transcriptional regulator mRNA	\neg	$\neg \tau$	U34279 Human uroguanylin mRNA, complete cds.	AA287021 2557c03.s1 Soares NbHTGBC Homo sapiens cDNA clone 701572 3'	yb47a01.s1 Homo sapiens cDNA clone 74280 3 containing L1 T55226 repetitive element	yf56e10.s1 Homo sapiens cDNA clone 26129 3' similar to gb:X07173	K3/446 INTER-ALTHA-INTESIMINATION COMPLEX COMPONENTS	AA406180 zu65c08.s1 Soares testis NHT Homo sapiens cDNA clone 742862 3'	R09752 Unknown	R81530 yj02b10.r1 Homo sapiens cDNA clone 147547 5'.	132348 EST47211 Homo sapiens cDNA 3' end similar to None		W57810 340946 3'	zt47e12.s1 Soares ovary tumor NbHOT Homo sapiens CDNA clone	AA33832 112318 3	1	vg52g07.s1 Homo sapiens cDNA clone 36232 3' similar to gb:M33987	R46266 CARBONIC ANHYDRASE I	H98618 yx12a06.s1 Homo sapiens cDNA clone 261490 3.	2097h01.s1 Stratagene ovarian cancer (#937219) Homo sapiens cDNA	AAT/1703 clube 374603 5 AAT/1703 clube 261854 3"	האינות אלוו אלוו אלווים אלווים אלווים אלווים אלווים אלוו אלווים
-	<u>∞</u>	0	76	0	0	9	2	2	0			٠,	-	\downarrow		7	7	_				15	+	╀	_	_		+	\downarrow
-	7	0	17	0	0	2	25	7	12	의	3		╁╌	+		0	4	-	-	_		1	- ~	╀	0	7	_	+	4
-	9	9	5	0	0	0	9	7		0	0	4	-	+		0	0	┞	\vdash	_		+	,,,	+	_		-	+	\dashv
-	=	8	8	9 /	_	3	2	12	-	0		,	┼	-		3	3	H	-			+	2 5	+	31	8	-	+	\dashv
-	88	8	8	37	37	37	38	33	34	34	34	2	\	\dashv	<u> </u>	 	=	+	-			-	7	+		\vdash	-	\dashv	4
	H328308	H434907	H618121	H349706	H259108	H611050	H241323	H386390	H950457	H740629	H511670	921C02H	00170011			11610982	111047673						H38/054	H90931	H390158	H893564			
	71 CATGCCAAAGCTATA	22 CATGCGGGAGTCGG	71 CATGCCGTGGAGAG	74 CATGCCCGAAGCC	75 CATGATTTCAAGATG	76 CATGGCCAGTGGCT	77 CATGATGGTGGGGA	78 CATGCCTCCCCCT	19 CTAGTGGAAGTGAA	80 CATGGTCATCACCAC	81 CATGCTTATGGTCCC	CHOCOCOCOCOCOCOCOCOCOCOCOCOCOCOCOCOCOCO	82 CATGCTGGGCCTCTG					84 CA101111AC1CA1						% CATGACCTGGGGAGG			20 CATOLOGO A CATOLOGO		

		İ	l	Ì	ŀ		AND Susings Completing NALIBIT Home sanians
						AA02	ZK10e12.51 Societs, pregnant uterus rooms suprem control to the su
VO CATGGGAGGTGGGGC	H666539	30	9	~	32	22 M75	M75161 H.sapiens granulin mRNA, complete cds.
	H1003970	9	7	3	9	17 T30344	
CATGGTCTGGGGGAT	H752297	53	_	3	6	3 T60135	\neg
						T30403	gb U67963 HSU67963 Human lysophospholipase homolog (HU-K3)
			1	\dagger	\dagger		\top
	108414	20	~	0	∞	0 R23595	
92 CA IGITAACCCCTCC	1170411			T		┼	
						R69445	
				┞			yi84h01.s1 Homo sapiens cDNA clone 145969 3' similar to gb:D26129
						R79191	
							yj56c03.s1 Homo sapiens cDNA clone 152740 3' similar to gb: D26129
						R49965	
					\vdash		zv35h12.r1 Soares ovary tumor NbHOT Homo sapiens cDNA clone
							755687 5' similar to TR:G459890 G459890 OVEREXPRESSED IN
	H231029	78	~	~	4	6 AA41	AA410947 TESTICULAR TUMORS
93 CATGATGACGCICAC	11231027		1			T	20 yj40c11.r1 Homo sapiens cDNA clone 151220 5'.
			T	T	\vdash		1
						_	586718 5' similar to TR:G459890 G459890 OVEREXPRESSED IN
						AA13	AA130551 TESTICULAR TUMORS.
							zd33c10.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone
	H286420	28	Š	0	~	4 W68230	
CATOCACCIOICATO							yp90a02.s1 Homo sapiens cDNA clone 194666 3' similar to contains Alu
						R89822	_
						- 	2k69e08.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
					-	AA05	AA053322 488102 3' similar to contains element MER6 repetitive element
STOCATOCATO	H578824	27	-	-	24	17 V00594	594 Human mRNA for metallothionein from cadmium-treated cells
		,	-	,	-	K H43742	yp21d05.r1 Homo sapiens cDNA clone 188073 5' similar to gb:J05021
% CATGCTTAGAGGGG1	H310163	3 6	- -	1	\ \ -	╁	_
97 CATGATGGCCCATAC	H238923	33	-	٦	-	╀	V00497 Human messenger RNA for beta-globin.
98 CATGGCAAGAAGTG	H391664	7	-	,	1	4	

TT VOLUCTA COTOTA TT	H810468	27	5	1	=	12 X65	614 H.s	X65614 H.sapiens mRNA for calcium-binding protein \$100P.
100 CATGLEGE ACT	H233106	28	0	7	0	2		-
					-		E	emb Z69881 HSSERCA3M H.sapiens mRNA for adenosine
101 CATGTTCTGTAGCCC	H1014566	25	\$	0	4	0	П	triphosphatase, calcium
Invication	H388582	24	_	2	_	3 T99568		ye65c02.r1 Homo sapiens cDNA clone 122594 5'.
72						T87	T87539 yd8	yd89f09.s1 Homo sapiens cDNA clone 115433 31.
						_	qg	gb AA347726 AA347726 EST54132 Fetal heart II Homo sapiens cDNA
103 CATGTATGATGAGCA	H844682	23	4	0		0	5' e	end similar to transmembrane secretory component
104 CATGCTGGCAAAGGT	HS00747	23	0	0	0	0	$\overline{}$	
105 CATGCTTGATTCCCA	H517078	23	4	4	11	7 1.42		Homo sapiens bone-derived growth factor (BPGF-1) m
INFICATGETTGACATACC	HS16402	22	0	0	7	2 X68	X68277 H.s	phase
							丑	Human N-benzoyl-L-tyrosyl-p-amino-benzoic acid hydrolase
102 CATGGCTGGCACATT	H649492	22	S	0	0	0 M82		alpha subunit (PPH alpha) mRNA, complete cds
108 CATGTCTGAATTATG	H909556	71	-	-	_	1 X16	X16354 Hur	Human mRNA for transmembrane carcinoembryonic antigen (CEA)
					-		H.S	H.sapiens mRNA for Gal-beta(1-3/1-4)GlcNAcalpha-2,3-
TOAGGAAGACT	H657554	21	_	_		3 X74	X74570 sial	sialyttransferase
200000000000000000000000000000000000000				\vdash			yo ₄	yo45d01.s1 Homo sapiens cDNA clone 180865 3' similar to contains
AUDULTTUUTUUT AUJUIT	H646998	20	7	0	_	0 R87	R87768 PTI	PTRS repetitive element
				\vdash		_	y03	yo36g07.s1 Homo sapiens cDNA clone 180060 3' similar to contains
						R85	R85880 PTI	PTRS repetitive element
THE CATE AATCTEGE AC	1114245	2	~	0	4	3 L20	L20826 Hun	Human I-plastin mRNA, complete cds.
TI CATOMANICIONIO	H802708	6	7	0	-	7 ZS0	ZS0751 HS	HSB4BMR H.sapiens mRNA for B4B
				T	-	170	U77085 Hui	Human epithelial membrane protein (CL-20) mRNA, complete cds
						Y07	Y07909 HS	HSPAPR H.sapiens mRNA for Progression Associated Protein
270700000000000000000000000000000000000	H764570	∞	-	-	∞	2 R48	R48529 yj6	yj64g10.r1 Homo sapiens cDNA clone 153570 S'.
					-		ES	EST10a24 Clontech adult human fat cell library HL1108A Homo
LIACATGTTATGGTGTGA	H998127	17	0	0	_	727 0	T27534 sap	sapiens cDNA clone 10a24.
11 SCATEGORA ACAGO	H663571	=	-	2	4	0 T86	T86124 yd8	yd84b04.s1 Homo sapiens cDNA clone 114895 3'.
						_	02	zo15g05.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
						AAI	AA131008 587000 3'	000 3'
				T		R49	R49945 yjS	yj58g11.s1 Homo sapiens cDNA clone 152996 3'.
						T57	T57044 ya8	ya84h01.s1 Homo sapiens cDNA clone 68401 3".
116 CATGCCAACACCAGC	H328787	17	-	0	0	0		
TOATGAGGTGACTGGG	HI78299	13	0	0	0	0		
LISCATGGCCATCCTCCA	H609654	91	0	0	0	0	gp	gbR73013/R73013 yj94a09.rl Homo sapiens cDNA clone 156376 5.
10 01								

			-	ļ		-	MKODIZ	MACON 13 Human quanine nucleotide-binding regulatory protein
19 CATGTTTCTCGTCGC	H1039799	<u>-</u>	-	키	3	,	NIOSOIN	House East of the Control of the Con
CT CCTC TO	H860776	~		_	_	·		Unknown
120 CATOLCAGAOCOCTO	200		1	T	1			yv72h06.s1 Soares fetal liver spleen INFLS Homo sapiens
								cDNA clone 248315 3' similar to contains element PTR7 repetitive
	H1006014	7	_	0	0	2	N58523	element
21 CATOTACOGTOTOGO	H814011	4	-	0	0	0		Unknown
CATGCTCAGACTIG	H477216	4	0	_	4	2		Unknown
CATGGGACTAAATGA	H662543	=	-	0	_	0	M29540	M29540 Human carcinoembryonic antigen mKNA (CEA), complete cos.
								HUMGS04154 Human colon 3 directed Mbol cUNA, HUMUS04134,
TT & DOUBLE DOUBLE	H653988	12	0	0	0	-	D25786	clone cm0215.
CATOOCITOOCAT			T	T				yc36e02.rl Homo sapiens cDNA clone 82778 5' sımılar to gb:LU / 165
							T73613	LIVER CARBOXYLESTERASE PRECURSOR
	1186138	2	-	0	0	-		Unknown
126 CATGACCCAACTUCC	1001001	2	-	-	1	2		pb/T95615 T95615 ye40e03.s1 Homo sapiens cDNA clone 120220 31.
127 CATGCTGAACCTCCC	1491094	*	1	,	1	T		zr19b11.s1 Stratagene NT2 neuronal precursor 937230 Homo sapiens
	2011201	=	_	_	7	0	AA226797	AA226797 cDNA clone 663837 3'
128 CATGCAAGAGTI ICI	7011/7H		, T	,	1			zo97h01.s1 Stratagene NT2 neuronal precursor 937230 Homo sapiens
							AA218730	AA218730 cDNA clone 649969 3'
				T	T	Γ		vp57f10.r1 Homo sapiens cDNA clone 191563 S' similar to gb:M90657
A JOTO A COOTOOT A O	H743610	=	-	0	•	~	H38178	TUMOR-ASSOCIATED ANTIGEN L6 (HUMAN);.
Zy CATOOTCCOAOTOCA	111042445	E	c	6	0	0		Unknown
130 CATGTTTGGTTTCAC	H1042442			,	1			

cell lines compared to normal colon (78 genes) Transcripts decreased in only colon cancer

NC: Normal Colon

TU: Colon Primary Tumor

CL. Colon Cancer Cell Line PT. Pancreatic Primary Tumor PC. Pancreatic Cancer Cell Line

		72	1111	5	Z	PC	Accession	Gene Name
	I ag Inumoer	2	2	3 :		233		H sanjens mitochondrial EST sequence (1-t-12)
CATGCACCTAATTGG	H285759	612	2	=			- 1	II emisse natial cDNA comence: clone c-39e04.
CATGATTTGAGAAGC	H260227	603	566	22	\$	2	П	n. Sapiens parinal edition of the contents (ARS) mRNA
CATGTGATTTCACTT	H933704	452	595	235	8	314	П	Tuman autonomously reputcating sequence (1992)
CATCTTCATACACCT	H1002566	444	357	114	8	ᅙ	$\neg T$	H.sapiens mitochondrial ES1 sequence (001114)
CACTO	H115432	385	402	223	278	132	X51525	Human cortex mKNA containing an And repetitive cicinent
CATOCCACIOCACIO	H114966	369	446	12	26	191	F16402	H.sapiens mitochondrial ESI sequence (141-20)
CAIGACIAACACCI	H291282	293	527	78	14	83	U09500	Human mitochondrion cytochrome b gene, partial cus
CATGAAACATTCTC	H1272	200	691	86	17	223	\Box	H.sapiens mitochondrial ESI sequence (101-03)
CATGCTCATAAGGAA	H478249	184	127	70	21	75	$\neg \tau$	H.sapiens mitochondrial ES1 sequence (1-1-07)
CATGTCGAAGCCCCC	H885334	147	183	8	\$	22	\neg	H.sapiens mitochondral E.S.1 sequence (VZZ117)
CATGACGCAGGGAGA	H103075	145	160	۳	69	4	Т	yla /aug.s. Homo sapiens Condition 151002 5.
1) CATGTTGGCCAGGCT	H1025322	124	194	S	=	~	$\neg r$	H.Sapiens michae 101 Mino class 11 maister eds
CATGTTGGTGAAGGA	H1027595	86	106	1	≅	<u>6</u>	Т	Human inymosin peta-4 lintars, compare care (xe31)
CATGATCACGCCTC	H214616	16	186	1	4	49	_1	Human EST Overexpressed in painted the biggs (ASS)
15 CATGTGCCTGCACCA	H941638	<i>L</i> 9	48	25	75	×	- 1	Human mKNA for cysteine proteinase minorio precurso
CATGAGACCCACAC	H136465	8	121	28	24	2	$-\mathbf{r}$	Human tetal brain cDNA 3-end GEN-122003.
CATGAGTTTGTTAGT	H196339	8	33	11	13	\simeq	-	Human mKNA for adenocarcinoma-associated antigen
CATGGGACAACAG	H656389	2 8	41	4	=	3	\neg	Homo sapiens CD24 signal transducet mindra
CATGTGGTGTATGCA	H965434	53	27	9	20	5	Т	Human Tetal Oralin CDIVA 3 -clid OEN-002710.
CATOL A A TACAGET	H527436	49	35	01	100	36		Human cathepsin D mKNA, complete cos.
100001	91757H	49	37	21	12	15	U25801	Human Tax1 binding protein mRNA, partial cds.
CATGGIGGCICACGC	0033721	Y	74	<u>~</u>	23	15	U31215	Human metabotropic glutamate receptor 1 alpha
CATGGTGGTGCACAC	4/02/0/H		1	-	٧	-	S79597	tRNASer(UNC) [human, muscle, MERRF/MELAS overlap s
CATGGGGTTGGCTTG	H 704160	<u> </u>	3 5	1 2	, 5		T48809	vb05c03.r1 Homo sapiens cDNA clone 70276 5' contai
CATGGTGGCGGGTGC	H763567	7	7	-	3 3	, 5	Т	Human Plobin gene.
25 CATGTAGACTAGCAA	H821029	2	2	-	3	2	٦.	9

Human fetal brain cDNA 3'-end GEN-007C04.	2b91h11.s1 Soares parathyroid tumor NbHPA Homo sap	H.sapiens mitochondrial EST sequence (132-20) from skeletal	muscle	EST186995 HCC cell line (matastasis to liver in mouse) II Homo	AA315049 sapiens cDNA 5' end	H. sapiens partial cDNA sequence; clone A6A03; ver	yw53h01.s1 Homo sapiens cDNA clone 255985 3'.	Human MHC class I HLA-A2 gene, complete cds.	yf25f12.s1 Homo sapiens cDNA clone 127919 3'.	yi22c10.s1 Homo sapiens cDNA clone 158994 3'.	EST58371 Homo sapiens cDNA 3' end similar to None	H.sapiens mitochondrial EST sequence (129-09)	2154f10.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone	726187 3'	2131c11.rl Soares ovary tumor NbHOT Homo sapiens cDNA clone	AA292466 723956 5' similar to TR:G205858 G205858 RAT ORF	2b62d07.s1 Soares fetal lung NbHL19W Homo sapiens cDNA clone	308173 3' similar to PIR:A39484 A39484 androgen-withdrawal	apoptosis protein RVP1, prostatic - rat	zb19c06.s1 Homo sapiens cDNA clone 302506 3' similar to	PIR:A39484 A39484 androgen-withdrawal apoptosis protein RVP1,	prostatic - rat;	zk39d06.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA	clone 485195 3' similar to PIR: A39484 A39484 androgen-	withdrawal apoptosis protein RVP1	Human partial cDNA sequence with CCA repeat region	Human episialin variant A mRNA, 3' end.	Unknown	seq816 Homo sapiens cDNA clone b4HB3MA-COT8-HAP-Ft	H.sapiens mRNA for LI-cadherin.	Homo sapiens huntingtin (HD) gene, exon 66.	dbj C00470 C00470 HUMGS0007620, Human Gene Signature, 3:-	directed cDNA sequence.	lyy62g08.s1 Homo sapiens cDNA clone 278174 3'.
DS1017	W15552		F16326		AA315049	F01150	N29971	K02883	R09140	R76005	T33596	F16449		AA292959 726187 3'		AA292466			N92384			N80203			AA039323	U21468	M34088		T10098	X83228	L27415		C00470	N63531
2	=		6		2	36	7	2	~			2		7		7										2	11	0	4	2	7		~	
25	59		91		8	17	9	2	20			7		_		_										2	45	0	3	0	7		-	
=	٥		=		=	=	0	-	-			6		_		_										7	0	_	2	2	-		4	
144	372		170		13	<u>∞</u>	5	4	32			55		6		•										218	2	6	=	6	7		~	
38	37		37		33	33	22	32	32			82		28		76										78	25	24	24	22	21		21	
H641789	H687915		169669H		H261569	H294488	H386963	H132598	H489822			H609624		H610922		H956860										H175872	H387596	H188027	H353760	H2235	H607977		H167659	
24 CATGGCTAGGTTTAT	┰	+-	CATGGGGGTCAGGG	_	% CATGATTTTCTAAAA		_	\top	Т	_		34 CATGGGGATCCCTT	-		$\neg \neg$		200000000000000000000000000000000000000									33 CATGAGGGTGTTTC	_	┰		_	Т	\top	41 CATGAGGATGTGGG	\top

									zo80f04.s1 Stratagene ovarian cancer (#937219) Homo sapiens
								AA165679	AA 165679 cDNA clone 593215 3'
									zv40a02.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone
14	CATGTATAGTCCTCT	H838494	20	7	_	3	4	AA411012 756074 3'	756074 3'
									z192g08.s1 Stratagene colon (#937204) Homo sapiens cDNA clone
								AA133595 512126 3'	512126 3'
									zt56b12.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone
								AA292774 726335 3	726335 3'
Š	CATGGGTCCTCTCTT	H710520	2	1	7	7	2	R53216	yj73h02.rl Homo sapiens cDNA clone 154419 5' simil
T	CATGATGGGCTTGAT	H240121	61	4	0	3	3	D20113	Human HL60 3'directed Mbol cDNA, HUMGS01086, clone
${}^{-}$	CATGCTGCCCCCAT	H496981	6	~	0	ı	4		Unknown
	CATGITCICIACACA	H1013522	61	4	-	∞	2	U35048	Human TSC-22 protein mRNA, complete cds.
_	CATGAAGAAGCAGGG	H33355	<u>∞</u>	4	7	7	8	R81767	yj05g03.r1 Homo sapiens cDNA clone 147892 5'.
1	CATGAGTAGGTGGCC	H183018	<u>∞</u>	==	7	11	. 1	D51021	Human fetal brain cDNA 3'-end GEN-007D07.
	CATGACAGTGTGT	H77551	<u>∞</u>	~	~	0	8	D26146	Human DNA for putative protein kinase.
Т	CATGGGAAAGTGGT	H655547	∞	2	3	02	1	M11465	Human alpha-1-antitrypsin mRNA, complete cds.
	CATGAAGAAAGCTC	H32926	12	4	0	~	_	R78188	yi81g01.r1 Homo sapiens cDNA clone 145680 5'.
ı	CATGACACCCATCAC	H70965	=	4	0	0	0	M22406	Human intestinal mucin mRNA, partial cds, clone SM
	CATGAGATCCCAAGG	H144707	17	∞	0	0	0	T24507	EST082 Homo sapiens cDNA clone 3E6
_ [.									za63a11.s1 Homo sapiens cDNA clone 297212 3' similar to
				*		•		N79237	PIR:S49589 S49589 cortical granule lectin - African clawed frog ;.
								T31354	EST30893 Homo sapiens cDNA 5' end similar to None
\$	CATGAATAGTTTCCC	H52214	91	4	0	0	0	H54696	yq92e02.s1 Homo sapiens cDNA clone 203258 3' simil
	CATGCAGAAAGCATC	H295060	91	6	0	0	0	M22430	Human RASF-A PLA2 mRNA, complete cds.
	CATGGCTTTGCTTTG	H654976	9	4	7	∞	-	AA374631	EST86866 HSC172 cells I Homo sapiens cDNA 5' end
\neg					Γ				zn93g08.r1 Stratagene lung carcinoma 937218 Homo sapiens
								AA137163	AA137163 cDNA clone 565790 5'
						-			zk10f05.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA
								AA029320	AA029320 clone 470145 3'
.5	CATGIGGIGGIGGATIGA	H948543	~	2	0	-	0	D25681	Human colon 3'directed Mbol cDNA, HUMGS04047, clon
 -									2r72g02.s1 Soares NhHMPu S1 Homo sapiens cDNA clone 668978
								AA253331	3,
								H05110	yl75f07.s1 Homo sapiens cDNA clone 43778 3'.
3	CATGCCATCGTCCTT	H341720	5	∞	_	_	10		Unknown
	CATGGACACACTCAC	H529013	4	23	0	0	0	AA297150	AA297150 EST112734 Colon I Homo sapiens cDNA 5' end
_									

		zk01e10.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA	AA026974 clone 469290 3'	zu 12c12.r1 Soares testis NHT Homo sapiens cDNA clone 731638 5'	similar to gb:M61900 Human prostaglandin D synthase gene,	AA405031 complete cds. (HUMAN);	gblU66894 HSU66894 Human epithelium-restricted Ets protein ESX		Human epithelial-specific transcription factor ESE-1b (ESE-1)		Human colon 3'directed Mbol cDNA, HUMGS06772	Unknown	ze88g07.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone	AA071520 366108 3'	za90h10.s1 Soares fetal lung NbHL19W Homo sapiens cDNA clone		zn52h06.s1 Stratagene muscle 937209 Homo sapiens cDNA clone	AA086292 561851 3'	Human HepG2 3'-directed Mbol cDNA, clone a-35.	IB2474 Homo sapiens cDNA 3'end.			zh75f08.s1 Soares fetal liver spleen 1NFLS S1 Homo sapiens cDNA		H. sapiens partial cDNA sequence; clone c-29h08.		ya31a06.s5 Homo sapiens cDNA clone 62194 3' contains Alu	repetitive element,.	Unknown	Unknown	Unknown	Unknown	
M25629	H18836		AA02697			AA40503		U66894		U73843	D25996			AA07152		N90742		AA08629	D11499	T16031	T74426	17757N		W90388	F03786	U14631		T41121			Z58486		
0	7				_			•			0	_		0	L				0	0	-	7				0		3	0	0	0	0	
	~							6			_	_		7					0		9	_				0	_	3	2	0	0	0	
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4	2							6			٣	0		<u>~</u>					4	7		2				2	L	9	4	7	2	_	
=	2							13			2	=		. 22					12	2	2	2				2		=	=	Ξ	E	E	
H695406	11354776							H176584			H265232	H503809		H774358					H49304	H658173	H670333	H715099				H817952		H360008	H440966	H611590	H616862	H666014	
CATGGGGCTACGTCC	DELICATION OF THE PROPERTY OF							ATOACOTACTA	04 04 04 04 04 04 04 04 04 04 04 04 04 0		S CATGGAAATAAATTA	\neg	00 CA10C10L0	CATCOTTCA & TOOT	פין כאוססווכאזוכסכו				STUDENT A A TA A GLUTT	68 CATGGGAAGGTTTAC		1 CATOCOTOCOTOCO	7			CATGTACTGTACTTC		23 CATGCCTTGCACTC		т.		1	_1

2d42c12.s1 Soares fetal heart NoHH19W Homo sapiens CDIVA CIOITE	174226 11 11 0 0 0 W68073 343318 3' similar to contains Alu repetitive element;
	W68073
	0
	0
	0
	=
	11
	H874226
	78 CATGTCCCCGTTACA
- 1	2

BNSDOCID: <WO___9853319A2_I_>

Table 4 - Transcripts increased in pancreas_cancer .

SAGE Tags elevated only in Pancreatic Tumor NC Normal Colon The Colon Tumor CC Colon Cancer Cell Line PT Pancreatic Tumor PC: Pancreatic Cell Line

£	rancreatic Cell Lille				ŀ	ļ	t		Γ	
-	Tag Sequence	Tag Number	SC	2	Tu CC PT		2		٥	Cene Name
-	CATCABACCA	H9222	0 2	9	=	~	Ξ	Examples R38305		yh95b04.s1 Homo sapiens cDNA clone 13/455 3
-	100000000000000000000000000000000000000					\vdash				zk95b03.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
									AA126719	490541 3'
+					T	-				zk51c03.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
				_		_			AA044296	486340 3'
						╁				213308.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
									AA131586	503726 3'
+			I		T	\vdash				zo71h12.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
		H9408	_	~	7	21	m	Examples	Examples AA157983	592391 3'
7	AI GAAAGCAGTTTO				T	t	T			2154e04.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 726174
						_			AA292929	3-
+					T	\dagger				2078c07.s1 Stratagene pancreas (#937208) Homo 2078c07.s1 Stratagene
									AA159306	pancreas (#937208) Homo
+						\vdash	T		R54012	yj70h01.s1 Homo sapiens cDNA clone 154129 3'
+						+			T62936	yb99f08.s1 Homo sapiens cDNA clone 79335 3'
100	TOPECACA	H9898	0	P	0	0	=	Examples X52426	X52426	H. sapiens mRNA for cytokeratin 13
3 2	CATCABATCTEGGT	H13803	L	-	F	9	7	Examples X51698		H.sapiens spasmolytic polypeptide (SP) mRNA.
7 7	4 CATGAGATGGACAG	H14865	0	0	F	0	23	Examples N70419	N70419	za61d12.s1 Homo sapiens cDNA clone 297047 3'
3	AL GAME I SOURCE					\vdash			AA411599	zv16g01.r1 Soares NhHMPu S1 Homo sapiens cDNA clone 753840 5'
-						-			AA410508	zy 16p01.s1 Soares NhHMPu S1 Homo sapiens cDNA clone 753840 3'
+			I	\prod		\dagger	1		Т	2186g12.51 Stratagene colon (#937204) Homo sapiens cDNA clone 511558
- î	TOTTTORNORS	H21247	7		9	∞	13	Examples	Examples AA115723	3'
3	A10000000					\vdash	T			2019e04.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 587358
					_				AA132875	31
+						t	Γ			2044a06.s1 Stratagene endothelial cell 937223 Homo sapiens cDNA clone
					_				AA147677	589714 3'
_			_	_		4	1			

			-	-			200,004	zq81h12.s1 Stratagene hNT neuron (#937233) Homo sapiens cDNA clone
	0070011	+	-	12	17	Examples R41318	AA200883	04607113
CATGAACTCTTGAAG	H30069	+		1		Condition	T35270	EST82235 Homo sapiens cDNA 3' end similar to None
			_				AA412071	2(65h12.s1 Soares testis NHT Homo sapiens cDNA clone 727271 3'
A CONTRACTOR A	H31221	1-	6	9	130	Examples N63154		yz37f12.s1 Homo sapiens cDNA clone 285263 3'
		+	\vdash					yc81h04.s1 Homo sapiens cDNA clone 22603 3'
		-	\vdash				AA150720	2146f04.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone 5049
							AA045773	z168b12.s1 Stratagene colon (#937204) Homo sapiens
UCATGAACTTGGCCAT	H32405	6	0	0	=	Examples X07819	X07819	Human pump-1 mRNA homolog. to metalloproteinase,
		-	-	L			L22523	Human matrilysin gene, exon 5
11) CATGAAGATCCCCGC	H36183	~	<u> </u>	14 12	23	Examples R72650	R72650	yj95e05.s1 Homo sapiens cDNA clone 156512 3'
		-	-					Area con a constant the second
								2030c02.51 Source redainted in the HTDLIP of Them Saprems Court Court
							W70287	DIVALENT CATION TOLERANCE PROTEIN CUTA
		+	╀					yj95e05.s1 Homo sapiens cDNA clone 156512 3' similar to
								SP.CYCY_ECOLI P36654 C-TYPE CYTOCHROME BIOGENESIS
			_				R72650	PROTEIN CYCY
								Control of Control of the Control of
								Zpotatijsi Sudaggije endomental cen 95/225 mono sapicijs CDNA cionic 624668 31 similar io SW-CITA FCON 1936654 PERIPI ASMIC
							AA181976	DIVALENT CATION TOLERANCE PROTEIN CUTA
		\vdash	-					Human phosphotyrosine independent ligand p62 for tthe Lck SH2 domain
H CATGAAGGGAGGGTC	H43180	9	<u></u>	8 15	41	Examples U46751	U46751	mRNA, complete cds
CATCAAGTTGCTATT	H48756	2	6	18	27	Examples 103077	J03077	Human co-beta glucosidase (proactivator) mPNA
		-	-	L			M86181	Human prosaposin (PSAP) gene
		\vdash		_			D00422	Human sphingolipid activator proteins, mRNA
		\vdash	-				103015	Homo sapiens sphingolipid activator protein 1 mRNA
		┢	\vdash				M60255	Human mutant cerebroside sulfate activator protein
LICATGAATGAAAAAA	H57345	0	-	5 2	10	No Match		
LICATGACAAACTGTGG	H66031	-	4 24	4 5	99	Examples N22375		yw37d01.s1 Homo sapiens cDNA clone 254401 3'
		-					A A 084643	2n20e01.s1 Stratagene neuroepithelium NT2RAMI 937234 Homo sapiens [,
		\dashv	\dashv	4			7	

				-			
					····-	AA279290	2584a06.51 Soares NbHTGBC Homo sapiens cDNA clone 704146 3'
						Т	zfi 2002.81 Soarcs fetal heart NbtH19W Homo sapiens cDNA clone
				-		23	376682.3
15 CATGACAACTCAATA	H67396	2 7	7 16	3	Examples Z58016		H. sapiens CpG DNA, clone 26c/,
					·		2029c02 s1 Stratagene colon (#937204) Homo sapiens cDNA clone 588290
			-			AA151668	3' similar to SW:BI3_MOUSE P28662 BRAIN PROTEIN 13
		+				П	za07e06.r1 Soares melanocyte 2NbHM Homo sapiens cDNA clone 291874
	•					W02958	.\$
		-	-	\dagger			zo70e05.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
	H71151	-	0 2	- 4	Examples	Examples AA1556464	592256 3'
In CATGACACCCI 61 6C		+	L	-		_	2c90h09.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone
					•	AA025673	366305 3'
		+	-	-		N70895	za89h12.s1 Homo sapiens cDNA clone 299783 3'
	P6058H	~	5	4	Examples X02491		Human interferon-inducible mRNA (cDNA 9-27): membrane
/ CATGACCATTGGALI	77,001	_	1_	+			Human interferon-inducible protein 9-27 mRNA
		+	1	1		X84958	H.sapiens mRNA for interferon-induced 17kDa membra
	חסטטעט	4	2	-	Examples X56841		H.sapiens HLA-E gene.
SCATGACCCI II AACA	OCOCK!!	+	1_	\dagger			H.sapiens mRNA for HLA-E heavy chain (exons 4 - 7)
	97219H	49 22	45 70	ヌ	Examples M21186		Human neutrophil cytochrome b light chain p22A
ly CAT GACCGCCG1GG1	_1_	+		\vdash			Human p22-phox (CYBA) gene, exons 3 and 4
	H97158	6	787	12	Examples D00244		Human Pro-urokinase gene,
20 CATGACCT GIGACCA		1	1	-			Human urokinase gene, 3' end
		F	F	\vdash		M15476	Human pro-urokinase mRNA, complete cds
				-		X02419	Human uPA gene for urokinase-plasminogen activator
	H103912	-	= 0	7	Examples L08835		Human myotonic dystrophy kinase (DM kinase) gene
21 CATGACGCCCTGCTC	71/20111	,		+			Homo sapiens myotonin protein kinase (DM) mRNA
	0322117	4	2	2	Examples H44451		yo75f06.s1 Homo sapiens cDNA clone 183779 3'
22 CATGACGTGGTGATG	DOCCI III		上	+			zo42107.s1 Stratagene endothelial cell 937223 Homo sapiens cDNA clone
-							589573 3' similar to SW:L10K_RAT Q05310 LEYDIG CELL TUMOR 10
						AA157329	KD PROTEIN
	-		-	-			2c32g06.s1 Soares senescent fibroblasts NbHSF Homo sapiens cDNA clone
						W46455	KD PROTEIN
	-	1		1]-	

.. . .

	1110183	0	3 21	3	Examples M92357		Homo sapiens B94 protein mRNA, complete cds.
2) CATGACTCAGCCCGG	n 17362						
	H123521	0	0 53	3 22	Examples X64875		H.sapiens mRNA for insulin-like growth factor binding protein
לין ראו פארו פאפניסטפ			\vdash				Human growth hormone-dependent insulin-like growth factor binding
						M31159	protein 3
			-			M35878	Human insulin-like growth factor-binding protein-3
			\vdash			SS6205	insulin-like growth factor binding protein 3 (3' region)
	H174764	10	0 22	9	Examples U65932		Human extracellular matrix protein 1 (ECM1) mRNA
25 CATGACTGCCCGC16	וווייייייי	+					Human extracellular matrix protein 1 (ECM1) gene, exon 9
		+	+	_			zo03f09.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 566633
	H126208	4	6	2 22		Examples AA148916	3,
26 CATGACTGTATTTIC	00707111	L	1_				zo12a11.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 586652
						AA129137	31.
		+	\vdash				2185g09.s1 Stratagene colon (#937204) Homo sapiens CLINA Clone 311430
						AA115437	3'
		1	+	L			z187e07.s1 Stratagene colon (#937204) Homo sapiens cDINA clone 311020
						AA126967	3,
	30007111	1-	-	3 16	Examples R24613		yh36c03.r1 Homo sapiens cDNA clone 131812
27 CATGAGCACTGCAGC	H14939	1	\perp				vp05c05.r1 Homo sapiens cDNA clone 186560 5'
28 CATGAGCAGGAGCGT	H150055					Γ	H saniens ckshs2 mRNA for Cks1 protein homologue
29 CATGAGCTGTATTCT	H162622	7	╛			T	245007 s1 Spares pregnant uterus NbHPU Homo sapiens cDNA clone
	2777	-		10		Examples AA044081	486300 3'
10 TATGAGGATGACCCC	H16/440	7	1.	1		Τ	2K50g07,r1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
							486300 S' similar to PR; A40533 A40533 cAMP-dependent protein kinase
						AA044211	major membrane substrate
	1178170	4	0	60	Examples X14787	X14787	Class A, Human mRNA for thrombospondin.
il cardadererrear	1178603		1			R27738	yh64f11.s1 Homo sapiens cDNA clone 134541 3'
STOCKE STOCKER	Cook	L	+	_			yj22fi2.s1 Homo sapiens cDNA clone 149519 3' similar to SP. ZK637.5
						H00276	CE00436 ARSA
			+	-			zm19d07.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
	H183787	3	=	15 73		Examples AA076235	526093 3'
13 CATGAGTATC I GGGA		1_	\vdash			H13159	yj16c04.s1 Homo sapiens cDNA clone 148902 3'
	1.		+	-			zo71e11.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
						AA146632	592364 3'
	U200740	0	ļ-	81	9 Examples	Examples X80062	H.sapiens SA mRNA.
34 CATGATACTTTAALI	2112711			-		U01691	Human annexin V (ANX5) gene
			}	-			

			\mid	-			V12464	Himan mBNA for vascular anticoamilant
			\dashv	4			716474	ווחוושו וודי זה הספיר שייים משפח שייים ווחוושו וודי זה הספיר שייים ווחוושו וודי זה הספיר שייים ווחוושו וודי הסייר הייים ווחוושו וודי הייים וודים וודי הייים וודים וודי הייים וודים וודי הייים וו
				-			M18366	Human placental anticoagulant protein (PAP) mKNA
			_				M21731	Human lipocortin-V mRNA, complete cds
			-	_			103745	Human endonexin II mRNA, complete cds
			\vdash	\vdash	_			GAMMA-INTERFERON-INDUCIBLE PROTEIN IP-30 PRECURSOR
CATGATCAAGAATCC	H213518	7	_	5 25		1 Examples 103909	103909	(HUMAN)
			-	\vdash				EST97384 Thymus II Homo sapiens cDNA 3' end similar to interferon,
							aa383911	gamma transducer 1
GOTGATCAAGGGTGT	H213679	12	9	25 12	2 156	5 Examples U09953	U09953	Human ribosomal protein L9 mRNA
							U21138	Human ribosomal protein L9 mRNA, complete cds
							17631	Umman mDNA for human homologue of rat ribosomal protein
			+	4			1557.0	יייייייייייייייייייייייייייייייייייייי
CATGATCAAGTTCGA	H213751	0	. 7	· ·	3 10		Examples AA063259	zm03a05.s1 Stratagene comeal stroma (#93/222) Homo sapiens cDNA clone 513008 3'
#DDBDBDDT#BT#D 81	H219750	91	7	14 12	40	Examples L42856	L42856	RNA polymerase II transcription factor SIII p18 subunit mRNA
10 CATCATCA ACTOC	H229502	1	0	0 17		4 Examples 259242	259242	H.sapiens CpG DNA, clone 13a10, reverse read cpg1
			\vdash	-				
	H235631	,	-		3 22	Examples 225820	225820	H.sapiens mRNA for mitochondrial dodecenoyl-CoA dehydrogenase
THE CALL GALL GEOGRAPHICS		1					L24774	Homo sapiens delta3, delta2-CoA-isomerase mRNA
	H243676	0	-	 	14	Examples M84711	M84711	40S RIBOSOMAL PROTEIN S3A (HUMAN)
1) CATGATGTCTTTTCT	H243710	1	7	1 14			M62403	Human insulin-like growth factor binding protein 4
200000000000000000000000000000000000000			\vdash	-				Human insulin-like growth factor binding protein-4 (IGFBP4) gene,
							U20982	promoter and complete cds
11 CATGATGTGTAACGA	H244487	0	4	5 44	t 94	1 Examples 233457	233457	H.sapiens mts1 gene.
				_			M80563	Human CAPL protein mRNA, complete cds
11 CATGCAACTTAAAGC	H270083	0		2 10		1 Examples N23207	N23207	yx70b09.s1 Homo sapiens cDNA clone 267065 3' similar to gb:L12350 THROMBOSPONDIN 2 PRECURSOR (HUMAN)
								2(25e11.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 714188
14 CATGEACCTGTGTCCTT	H286424	-	4	2 10		1 Examples	Examples AA285023	3' similar to gb:M33680 CD81 ANTIGEN (HUMAN)
			\vdash	-	_		M33680	CD81 antigen
INCATCCACTCAATAAA	H291889	0	0	7	3 19	9 Examples D78203	D78203	Neurosin
			\vdash	L			U62801	protease M

17 CATGCAGCCTGGGGC	H300971	0	- 0	0	10	Examples	Examples AA149942	2068d04.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone 592039 3' similar to TR:E218488 E218488 TRYPTASE
								zp66b09.r1 Stratagene endothelial cell 937223 Homo sapiens cDNA clone 625145 5' similar to gb:M16937 HOMEOBOX PROTEIN HOX-B7
18 CATGCAGCGCGCCT	H301462	4	11 12	2 10	21	Examples	23	(HUMAN); contains element MER22 repetitive element
				Ц			M16937	Homeobox protein HOX-B7
JACATGCAGGTTGTCCT	H307126	0	0	4 0	2	No Match		VIVE. 0.5
SUCATGOAGTOTOTOAA	H309109	7	9	6 2	17	Examples U14972		Human ribosomal protein S10 mKNA
	H316857	0	<u>س</u>	3	13	Examples U27293		Human leukotriene A4 hydrolase gene
TAISCAICCES SAC		+	-	L			103459	Human leukotriene A-4 hydrolase mRNA, complete cds
		+	+					Human leukotriene A-4 hydrolase mRNA, complete cds
EHO JEU JEE ACCE	H375080	6	2	5 13	~	Examples X82434		H. sapiens mRNA for emerin
Statiscallecterit	H333138	1	17	7 18	7	Examples M88338		Human serum constituent protein (MSE55) mRNA
SI CATGCCACCCCACC	909060	۲	12	7 22	28	Examples U14971		Human ribosomal protein S9 mRNA
A CATGCCAGTGCCCG	H344031	1	1	1		Examples L01697	L01697	Homo sapiens alpha-1 type XV collagen mRNA
SSCATGCCATTTTCIGG	H344691	, =	1 00	18	1_	Examples X54079	X54079	Human mRNA for heat shock protein HSP27.
\$6 CATGCCCAAGCTAGC	17011	+		上			Z23090	H.sapiens mRNA for 28 kDa heat shock protein
		+	+	1			X16477	Human mRNA fragment for estrogen-regulated 24k protein
		+	+	-			S74571	estrogen receptor-related protein=27-kda heat shock protein
	11247400	ڄ	15 43	10	2	Examples X69392	X69392	H.sapiens mRNA for ribosomal protein L26.
\$7 CATGCCCATCCGAAA	H24/402	1	1	L_			L07287	Human ribosomal protein L26 (RPL26) gene
4 7 7 6 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7 7	H150099	-	+	6 14	25	Examples U40434	U40434	Human mesothelin or CAK1 antigen precursor mRNA
CAT GUCUCUI GUAGA		+	+	-				Human mRNA for pre-pro-megakaryocyte potentiating factor, complete
							D49441	cds.
E 4 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	HISTORI	6	6	8	Ξ	Examples U12819	U12819	Human p16-INK4 (p16) gene
SO CATGCCCCCAIAGAI	in Control	+	L	1			U38945	Human hypothetical 18.1 kDa protein (CDKN2A) mRNA
		+	+	1				MTS1=multiple tumor suppressor 1/cyclin-dependent kinase 4 inhibitor
							S69804	910
		+	+	1			S69822	CDK41=cyclin-dependent kinase 4 inhibitor
		\dagger	+	\downarrow				tumor suppressor gene, P16/MTS1/CDKN2=cell cycle cycle negative
							S78535	regulator beta form
	79872FH	~	~	5 14	34	Examples 247319	247319	H.sapiens mRNA for expressed sequence tag (clone 21fi7119)
(a) CATGCCTCCTGGGG	100/001	5	1	ı	1	1		

							AA398406	2160h12.s1 Soares testis NHT Homo sapiens cDNA clone 726791 3'
	H170034	4	-	4	6	Examples U21049		Human DD96 mRNA
CATGCCGCCCTACC	H387925	<u> </u>	7	8	8	Examples X03212		KERATIN, TYPE II CYTOSKELETAL 7
02 CA1 0C 1 1 0G 1 CC 0C		-					A A 187637	2p/3f01.s1 Stratagene HeLa cell \$3 93 /210 Holilo Sapiciis Colino (25849 31
		-		+	-			zp35g11.s1 Stratagene muscle 937209 Homo sapiens cDNA clone 611492
63 CATGCCTTTGAACAG	H392709	<u>~</u>	ब	7	2	Examples AA1 /043 /	Т	2013611 s1 Stratagene muscle 937209 Homo sapiens cDNA clone 611468
			-				AA176541	3' similar to TR: G663269 G663269 BOLA.
	11415944	12	45	150	1	Examples	Г	Human interferon-inducible mRNA fragment
64 CATGCGCCGACGATG	H475429	1_		9	2	Examples T53402		ya88g05.s1 Homo sapiens cDNA clone 68792 3'
6) CATGCTCAACAGCAA		1_	1	-	\vdash			and a MA aminos amolt Mattheway of the same
							13/20403	2d47g08.s1 Soares retaineart Noriri19 w norm seprem scrive scrive and 243818 31 einitar to PIR-524168 S24168 hypothetical protein - human
		-	1	\dashv	+	-	21001	Umman mBNA for I DI -recentor related protein
CALCATGCTCAACCCCCC	H475478	-	7	티	=	Examples A13910	113910	Tullian mach to the foreign
CA CATECTERGAAACTG	H493576	2	-	∞	<u></u>	Examples X80335	X80335	H. Sapiens (24) reminin 11 pseudogene.
COLLIE AGEORAGES	H494454	-	4	21	13	Examples X04828	K04828	Human mking for U(1) proteint appliars accuming
ASSOCIATION OF ASSOCI	H498887	16 30	0 28	30	4	Examples U14966	J14966	Human noosomal protein L3 inches
AGEORCECOECOECOECOECOECOECOECOECOECOECOECOECO	H499247	-	3 4	13	13	Examples T90665	190665	yd41g08.51 Homo sapiens curin cione 1100+0 3
/0 CATGCTGCTGAGTGA		+	L	\vdash	\vdash			EST43791 Fetal brain I Homo sapiens CUNA 3 end similar to steroid
							AA338799	hormone receptor hERR1
		+	1	\dagger	\vdash		H97236	yv98b06.s1 Homo sapiens cDNA clone 250739 3'
	H501337	-	4	0	2	Examples C14084	C14084	Human fetal brain cDNA 3'-end GEN-018D10
71 CATGCTGGCGCCGA1	H513181		23 36	E	5	Examples D00017	D00017	Human lipocortin II mRNA
72 CATGCTTCCAGCTAA	H\$14022	1	1	8	-	Examples Z19574	219574	H.sapiens gene for cytokeratin 17.
73 CATGCTTCCTTGCCT	22011011	+	┸	+	\dagger		X62571	H.sapiens mRNA for keratin-related protein
		+	L	\vdash			X05803	Human radiated keratinocyte mRNA 266
	H522198	-	7	19	4	Examples X79067	X79067	H.sapiens ERF-1 mRNA 3' end.
71 CATGCTTTCTTCCT	US24280	1_	14 21	18	37	Examples X51779	X51779	Human mRNA containing an Alu repeat
75 CATGGAAAAAAAA	C0747CU	1	1_	+			X82240	H.sapiens mRNA for Tcell leukemia/lymphoma 1
6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	U525348	4	7 14		22	Examples V00572	V00572	Human mRNA encoding phosphoglycerate kinase.
76 CATGGAAACAAGATG	05007011	+		+	+		D29018	Human keratinocyte cDNA, clone 001
		+	-	\dagger	T		L00160	Human phosphoglycerate kinase (pgk) mRNA
-	11617436	9	35 10	100	120	Examples X05344	X05344	Human mRNA for cathepsin D
77 CATGGAAATACAGTT	004/701	- 1						44

		-		-	-		MI 1223	Hinman cathensin D mRNA, complete cds
0		-	1	†	+			442 ma et Homo eaniene CDNA clone 110909 3' similar to SP. R151.9
SASTASTA A A SOTA S.	H527929	4	- <u>-</u>	7	56	Examples T90296		CE00827
							142	EST23523 Adipose tissue, brown Homo sapiens cDNA 3' end
		+		\dagger	╁		П	zp64f07.s1 Stratagene endothelial cell 937223 Homo sapiens cDNA clone
	H513436		7 16	9	78	Examples AA181811		624997 3'
CATGGAAGATGTGTG	2000	┸	1	\dagger	1			2106c06.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
				-			08	491530 3' similar to WP:ZK652.2 CE00448
	H540621	9	3 10	6	82	Examples L21950		Human peripheral benzodiazepine receptor related mRNA
CALGGAATITION OF				\vdash	\vdash			Human peripheral benzodiazepine receptor (hpbs) mRNA
	H540673	-	2 10	٣	17	No Match		
CALGGACACCACCACCACCACCACCACCACCACCACCACCACC	H545152		0	E	7	Examples U19718		Human microfibril-associated glycoprotein (MFAP2).
EUCOCACORTO TA	H545430	0	0	2	<u>~</u>	Examples M75165		H.sapiens epithelial tropomyosin (TM1) mRNA
וויין האו פפארנאפפרני		-			\vdash	_		Human fibroblast muscle-type tropomyosin mRNA
		1		\dagger			M74817	Human tropomyosin-1 (TM-beta) mRNA, complete cds
	05077517	1,	0	19	2	Examples M74092	M74092	Human cyclin mRNA
S.I CATGGACCCCAAGGC	0124671	_[_	10	=	्रि	Examples L37033	Γ	Homo sapiens FK-506 binding protein homologue
S CATGGACCLIGCCI		1	١.		\dagger			zb37g02.s1 Soares parathyroid tumor NbHPA Homo sapiens cDNA clone
	H548062		0	13	_	Examples N90046		305810 3'
TOTAL CONTRACTOR OF THE CONTRA		-		\vdash				z106a10.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
							AA115048	491514 3'
99409090400#10	H551315	<u>س</u>	2	32	2	Examples M63193	M63193	Human platelet-derived endothelial cell growth factor
いっていっつつつつつこと () () () () () () () () () (H554876	-	3	0	4	Examples M61764	M61764	Human gamma-tubulin mRNA,
CALCACTOR CACTUTES	H559615	0	0	7	2	Examples D17793	D17793	Human mRNA (HA1753) for ORF
OF OF OR OWN OF THE PROPERTY O	H560056	0	8	32	=	Examples S68252	S68252	TIMP-1=metalloproteinase inhibitor
מוסיום וספאסאים ואיז וועי		\vdash		\vdash	\vdash		X02598	EPA glycoprotein (erythroid-potentiating activity)
		-		t	\vdash		X03124	tissue inhibitor of metalloproteinase 2
4) CATGGAGCAGGATGA	H561807	0	0	-	2	No Match		
	787786			4	13	Examples	Examples AA214523	2189c01.51 Soares NbHTGBC Homo sapiens cDNA clone 682848 3'
V2 CATGGAGGGAGIICC	20010	+	_	T			N30324	yw/5d01.s1 Homo sapiens cDNA clone 258049 3'
COROCOLUMN CORAC CO	H570787	0	0 2	-	2	Examples X70070	X70070	H.sapiens mRNA for neurotensin receptor.
CAI GGAGI CCGGAGC	959CL5H	6	3	0	2	Examples H57673	H57673	yr27a10.s1 Homo sapiens cDNA clone 206490 3'
94 CATGGAGTTATGLIG	lacos (CI)	1		1				

135766 1 Soares freal heart NOHHISW Home supters CDNA clone 135766 1 Soares freal heart NOHHISW Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 135766 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13576 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13576 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13576 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13576 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13576 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13576 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13576 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13576 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13576 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13769 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13769 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13769 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13769 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13769 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13769 1 Soares pregnant uters NOHIPU Home supters CDNA clone 13769 1 Soares										
H582913 3 2 2 19 Examples AA046631 H587800									*	ze12c08.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone 358766 3' similar to SW:YA94_SCHPO Q09783 HYPOTHETICAL 11.4 KD PROTEIN C13G6.04 IN CHROMOSOME I
H585913 3 5 2 2 19 Examples AA046631 R91942 R91945 R91945 R91942 R91945 R91945 R91945 R91945 R91945 R91942 R91945	10	TOCAGTTCGACCT	H572806	7		1	29	No Match		
H585913 3 5 2 2 19 Examples AA046651 H587800 1 0 5 1 12 Examples U60205 H587825 17 13 29 73 38 No Match H605956 2 10 8 3 55 Examples X60489 H611597 1 4 1 47 9 Examples X15256 H611891 8 5 2 44 3 Examples X13425 H613577 3 8 5 27 6 Examples AA053346 H655177 1 6 7 13 10 Examples M38259 H655177 1 6 7 13 10 Examples M38259 M60748	<u> </u>			-	-					zk72d06,s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
H587800 1 0 5 1 12 Examples U60205 H589825 17 13 29 73 38 No Match H605956 2 10 8 3 55 Examples X60489 X60656 H606471 0 0 0 12 1 Examples X15256 H611597 1 4 1 47 9 Examples X15256 H611597 1 4 1 47 9 Examples X15256 H611884 0 0 1 3 16 Examples AA054483 H61884 0 4 4 23 39 Examples X13425 H613357 3 8 5 27 6 Examples AA053346 H643707 12 29 24 35 35 Examples AA053346 H655177 1 6 7 13 10 Examples M38259 M60748	C.A.	TGGATTAAGTGAG	H585913	ᆔ			হা	Examples	Т	488363 3'
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H606471 0 0 12 1 Examples U08021 H611597 1 4 1 47 9 Examples X15256 H611597 1 4 1 47 9 Examples X15256 X14829 X14829 X14829 X14829 X14829 X14839 X14881 X14829 X14839 X14881 X14829 X14838 X14829 X14838 X14881 X14829 X14838 X14881 X14829 X14838 X14881 X14839 X14838 X13425 X13426 X134	<u></u>	ופפרעווועיייו		1	-		Г			H.sapiens mRNA for elongation factor 1-beta
H611597 1 4 1 47 9 Examples X15256 X14829 X14829 304456 S44881 S44881 H616224 0 0 1 3 16 Examples AA054483 H618841 0 4 4 23 39 Examples AA136985 H633577 3 8 5 27 6 Examples AA136985 H643707 12 29 24 35 35 Examples U43368 H655177 1 6 7 13 10 Examples U43368 H655177 1 6 7 13 10 Examples U43368 H655371 1 8 30 16 38 Examples M60748	1 6	400440000	H606471	-	l	l	-	Examples		
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H616224 0 0 1 3 16 Examples AA054483 H617891 8 5 2 44 3 Examples X13425 H618841 0 4 4 23 39 Examples X13425 H633577 3 8 5 27 6 Examples AA136985 H643707 12 29 24 35 35 Examples U43368 H655177 1 6 7 13 10 Examples U4368 H655361 11 8 30 16 38 Examples M60748	\downarrow								S44881	HI.14=beta-galactoside binding protein
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H633577 3 8 5 27 6 Examples AA136985 H643707 12 29 24 35 35 Examples U43368 H655177 1 6 7 13 10 Examples U43368 H655361 11 8 30 16 38 Examples M38259 M60748	5	TGGCCTACCCGAG	H618841	0			2	Examples	X13425	Human mkNA for panercane carcinoma market OA3321, o
H643707 12 29 24 35 35 Examples AA053346 H655177 1 6 7 13 10 Examples U43368 U52819 H655361 11 8 30 16 38 Examples M38259 M60748	5	TGGCGGGTGGAG	H633577			.1	9	Examples	AA136985	202003.31 States pregnam merus reen seprem
CATGGCTCAGCT LGGA H655171 1 10 Examples U43368 CATGGCAAAAAAAA H655361 11 8 30 16 38 Examples M38259			LK41707				35	Examples	AA053346	zl70h04.sl Stratagene colon (#937204) Homo sapiens cDNA clone 510007 3' similar to gb:Z21507 ELONGATION FACTOR 1-DELTA
CATGGGAAAAAAA H655361 11 8 30 16 38 Examples M38259 M60748	3 5	TGGCTCAGCTGGA	H655177	I		I	02	Examples	U43368	Human VEGF related factor isoform VRF186 precursor, 0
H655361 11 8 30 16 38 Examples M38259 M60748		200211110001		\vdash	-				US2819	Human vascular endothelial growth factor B 186
M60748	15	TGGGAAAAAAA	H655361	=	1 1		38	Examples	M38259	Human cytochrome c oxidase subunit VIb
) -			\vdash	-	L		,	M60748	Human histone H1 (H1F4) gene, complete cds

			-					
		\dagger	+	1			M73240	Human (clone SF2) hepatacyte growth factor (HGF)
	1455547	=	=	3 70		Examples X02920	X02920	Human mRNA for alpha 1-antitrypsin carboxyterminal, 0
1117 CAT GGGAAAAGT GG1		+		1_			X01683	Human mRNA for alpha 1-antitrypsin
			╀				V00496	Human messenger RNA for alpha-1-antitrypsin
		\vdash	┞				100067	Human alpha-1 antitrypsin gene, 3' end
		+	+					2122601.51 Soares pregnant uterus NbHPU Homo sapiens cDNA clone
	H658059	0	-	4	16		Examples AA127040	502633 3'
TO W. GOSANGOON		+	\vdash					2d86t06.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone
			-				W81387	347555 3'
		+	╀				H45477	yo72h08.s1 Homo sapiens cDNA clone 183519 3'
	H666943	16	12	9	32	Examples D26598	D26598	Human mRNA for proteasome subunit HsC10-II , 0
	79£7367	6	上	1			N74310	za78c01.s1 Homo sapiens cDNA clone 298656 3'
17. CAT 566A61616161		+	-				H92750	yi92e01.s1 Homo sapiens cDNA clone 231768 31
		-	_				T24084	sea2272 Homo sapiens cDNA clone ssb4HB3MA(extended-ft-6) 3'
	116711456	+	1-	2	21	Examples X17567	X17567	H. sapiens RNA for snRNP protein B
III)CATGGGATTGTCTGG	CC+110H	+				1_	M34081	Human small nuclear ribonucleoprotein particle SmB
	0557730	10	0	9	22	Examples M69054	M69054	Human insulin-like growth factor binding protein 6, 0
ווו נאבפפכנייו רארי	2001	· -	L		١		M62402	Human insulin-like growth factor binding protein 6
	H677753	10	-	7	4	Examples N74323	N74323	za78d08.s1 Homo sapiens cDNA clone 298671 3'
111111111111111111111111111111111111111		+	+	_			H46766	yo18f08.s1 Homo sapiens cDNA clone 178311 3'
-		\dagger	╀	1			H41102	yn88a08.s1 Homo sapiens cDNA clone 175478 3'
		+	╀	_				zm84b09.s1 Stratagene ovarian cancer (#937219) Homo sapiens cDNA
	H686815	0	-	3 13	22		Examples AA074777	clone 544601 3'
20121201202		\dagger						zm04a04.s1 Stratagene corneal stroma (#937222) Homo sapiens cDNA
							AA062735	clone 513102 3'
		+	+	-				zm63f12.s1 Stratagene fibroblast (#937212) Homo sapiens cDNA clone
							AA112905	530351 3'
LINCATORGRAPHICAGAT	H688713	22	1	0 6	72	No Match		
TA CATGGGGAGGGGTGG	H690863	7	m	1 16	7	No Match		
and	H690890	-	0	1 14	1			
TO CAI GGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGG	H693112	-	-	3 39	2	Examples V00523	, V00523	Human mRNA for histocompatibility antigen HLA-DR
CA16666CA16161		\dagger	\vdash	_			X00274	Human gene for HLA-DR alpha heavy chain a class II
		\dagger	+	ļ			1211121	Himan Hi A. DR sloha-chain mRNA

								11. 1. Lean chair man 2 Canb
							707001	numan ma-ur neavy chain gene, 3 main
上本で本でのでものしのHAT	H715401	F	4	01 01	14		Examples U18009	Human chromosome 17q21 mRNA clone LF113.
י ו פרפו פפפטיפטו		+	-	\downarrow			T33413	EST57778 Homo sapiens cDNA 3' end similar to None
		1	\vdash	-	_		T33339	EST57474 Homo sapiens cDNA 3' end similar to None
4 Double And	H778778	┢	 -	1 16	30]_	Examples M59911	Human integrin alpha-3 chain mRNA
TO CATEGORACITE CACCA	H728810	2	2	16 15			Examples X87689	H.sapiens mRNA for putative p64 CLCP protein
CAI GGI ACI GI GGGG	H737344		0	01		Example	Examples L12350	Human thrombospondin 2 (THBS2) mRNA
34 CA1661 CA661 CA	H752296	22	35.4	45 76	29		Examples D21261	Human mRNA (HA1756) for ORF
17) CA1 60 10 10 10 10 10 10 10 10 10 10 10 10 10		T	\vdash				D29543	Human keratinocyte cDNA, clone 686
	H752521	0	~	7 12	2		Examples H51290	yp07a05.s1 Homo sapiens cDNA clone 186704 3'
CA1661C1616A6A6		+	+	\vdash			N20338	yx44g12.s1 Homo sapiens cDNA clone 264646 3'
		t	╀	$oxed{\perp}$				zo76e09.s1 Stratagene pancreas (#937208) Homo sapiens cDNA clone
		_					AA158271	592840 3'
TO TO TO TO TO TO THE TOTAL OF	H752531	0	6	0	13	No Match		
CALCADIATE ACT	H753162	0	-	2	2	No Match		
CATGGT CIT GARGE	FCEP5CH	12	4	42 15	89		Examples X87373	Class C, H.sapiens RPS3a gene
129 CATGGTGAAGGCAGT	H754567	0	1~	-	2		Examples X08058	GLUTATHIONE S-TRANSFERASE P (HUMAN)
Life CA. GGI GAATGACGG	H760361	0	<u> </u>	-	1 25		Examples X51439	Human mRNA for serum amyloid A (SAA) protein
THE CATGOT GUEGARC	H761481	1	6	5			Examples U15008	Human SnRNP core protein Sm D2 mRNA
1.2 CAT GGT GCT GGAGAA	H762533	1=	-		6 34		Examples U62800	Cystatin M (CST6)
CALGGI GGAGGGCAC	F00897H	=	=	1	1_		Examples H46430	yo12h12.s1 Homo sapiens cDNA clone 177767 3'
CATGGTGGTACAGGA	COOCOUNT	†	1_	1 .				zf13a06.s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone
							AA047563	376786 3'
		+	+	┼-	L			zo13f02.s1 Stratagene colon (#937204) Homo sapiens cDNA clone 586779
							AA130701	31
SA COTTO A CHITCOTAC SE	H774629	0	7	_	13	3 Example	Examples X59288	H.sapiens gene for intercellular adhesion molecule
CALCAL TOOL TOOL CO.		T		-			M24283	Human major group rhinovirus receptor (HRV) mRNA
		1	╁	╀	_		103132	Human intercellular adhesion molecule-1 (ICAM-1)
		\dagger	\dagger	╀	_		M55100	Human cell surface glycoprotein P3.58 mRNA
U DE E E DE DE LE LA CASA E LA CASA	H781823	†	+	100	30 24		Examples K02765	Human complement component C3 mRNA, alpha and beta
WATER OFFICE FOR THE COLOR	H782013	178	011	14 340	_		Examples M17987	Human beta-2-microglobulin gene
CALCALIDITATION STORY	H782391	-	9	12	4 14		Examples D00760	Human mRNA for proteasome subunit HC3
CA A E E CO	99170ZH	_	-	9		12 Example	Examples X57025	INSULIN-LIKE GROWTH FACTOR IA PRECURSOR (HUMAN)
CATGTAAGGCTTAAC	H807703		7	15			E	
LIUCATGTAATTTTGGAA	100,40011	-	1	+		1		

	HR02793	F		+	-	No Match		
CATGLAALLIGGAL	100,001		٦	-	=	Fyamples X85373		H. sapiens mRNA for Sm protein G
CATGTACATTTTCAT	H806901	- 0	1	1	=	No Match		
12 CATGTACCCCGTACA	0/60001		7	2 2	1-	No Match		
LICATGTACCCTTCTAT	C76808H) -	2	: -	, ,	Evample: 102031		Human placental tissue factor (two forms) mRNA
11 CATGTAGGAAAGTAA	H82/43/	-	7	+	=	Evanipies 3	_	Human tissue factor mRNA, complete cds
		+	+	+	+	. ≥		Human tissue factor gene, complete cds
	717118H	49 61	12	68	130	Examples X64899		H. sapiens mRNA homologous to mouse P21 mRNA.
CATGTAGGTTGTCTA	ㅗ	1	1	+	-	×		Human mRNA for translationally controlled tumor protein
				\vdash				
			_			.1		Homo sapiens (clone 04) translationally controlled tumor protein
L. CATCTATATATCTC	H839672	0	٣	∞	91	Examples M98479		Human transglutaminase mRNA
TO THE THE THE COLUMN TO THE C	H851834	0	7	16	3	Examples D12149		Human HepG2 3'-directed Mbol cDNA, clone \$247
A A CO CA A CA C	<u> </u>	10 28	27	24	48	Examples X80909		H.sapiens alpha NAC mRNA
EAGUERA AUGEORIA	H868569	-	0	43	11	Examples X56134		Human mRNA for vimentin.
14 CA161CCAAA1CGA1		F	T	\vdash		2		H. sapiens vimentin gene
		1		+	\vdash	Σ	M14144	Human vimentin gene, complete cds
			T	+	+	Σ	M25246	Human vimentin (HuVim3) mRNA, 3' end
	H870310	0	†=	12	7	Examples N92906		zb57a08.s1 Homo sapiens cDNA clone 307670 3'
יייייייייייייייייייייייייייייייייייייי			T	╁	\vdash			
						<u>—</u>	T17488	NIB978 Normalized infant brain, Bento Soares Homo sapiens cDNA 3'end
		F	T	+	\vdash	A	AA349906	EST56900 Infant brain Homo sapiens cDNA 3' end
	HR71920	9 9	2	25	~	Examples X67016		H.sapiens mRNA for amphiglycan
CATCHICLETT		1_		\vdash				Human mRNA for ryudocan core protein
	H899060	2 5	2	+	69	Examples M77233		Human ribosomal protein S7 mRNA
CATGLCGICTITATO	H908858	1 5		46	6	Examples S48568		tissue inhibitor of metalloproteinase 2 (3'-end region)
TO TOTAL TOTAL	H916232	9	~	-	13	Examples N71680		yz93b03.s1 Homo sapiens cDNA clone 290573 3'
STATISTICATION OF A PARTICULAR	H916372	14 22	2	20	45	Examples X03083		Human lactate dehydrogenase-A gene
CAIGICI I GIOCOTO		-		-	-	×	X02152	Human mRNA for lactate dehydrogenase-A
	-			-	-	×	X02153	Human pseudogene for lactate dehydrogenase-A
	110201302	= = = = = = = = = =	۳	0	92	No Match		
156 CATGTGAAGTCACTG	7/207/11	1	I	+	+			
157 CATGTGAAGTTATAC	H920525	0 1	3	9	目	Examples X07979		CTGTGG, Class A, Human mRNA for fibronectin receptor beta subunit.
200000000000000000000000000000000000000				ĺ				

					ł	-			Liberor of Society are many when sub-HP11 Homo caniens CDNA clone
		1262223		~	. =	12	Examples AA027860	-	ZKOJIV. 31 Soares pregnam urotas rotas e como esperante de 180693 3º
.58	ISS CATGTGATGTCTGGT	H932/31	> -		-	: 2	Examples M25753	Т	G2/MITOTIC-SPECIFIC CYCLIN B1 (HUMAN)
2	CATGCCATCTGTA	0120001	+		1	1			yc22c04.s1 Homo sapiens cDNA clone 81414 3'
1			-	\sqcup		$ \cdot $		R67969	yi29g08.s1 Homo sapiens cDNA clone 140702 3'
1						-			2091f03.s1 Stratagene ovarian cancer (#937219) Homo sapiens cDNA
									clone 594269 3' similar to SW:NGAL_HUMAN P80188 NEUTROPHIL
9	[60] CATGTGCCTCAAAA	H939841	=	13 3	=	2	Examples AA169614	T	GELATINASE-ASSOCIATED LIPOCALIN FRECONSOR
									SW.NGAL_HUMAN PROISES NEUTROPHIL GELATINASE-
- <u>5</u>	IGI CATGTGCCCTCAGAA	H939849	3	4	=	2	Examples N79823		ASSOCIATED LIPOCALIN PRECURSOR
									zm90h04.s1 Stratagene ovarian cancer (#937219) Homo sapiens cDNA
	AS STOLLD CALL	H939851	13	31 10	25	83	Examples.	Examples AA075896	GELATINASE-ASSOCIATED LIPOCALIN PRECURSOR
	05540E000E0E40 671	H920392		_			No Match		MOLES and AMG animo II (1992)
2	CAleiecteichee		+	-		T			z181e07.s1 Stratagene colon (#937204) Homo sapiens civina cione 311044
	INSTRACTECCTIACTIT	H941856	0	3 1	7	2	Examples	Examples AA100279	31
	16.1 CATGTGCGCTGGCCC	H944038	2	5 2	17		No Match		STATE CONTRACTOR CONTRACTOR
									zkilda01.si Soares pregnani uterus indriro nomino sapiens contra cione
	STOTECTION	H949560	2	9 9	4	2	Examples	Examples AA029262	470088 31
						-		164701	yybbelu.si Soares ieläi livei spieeli iivi ksi lioliko sapielis veiva siolika
			\dashv	4		+		1074CN	2776603 s1 Stratagene NT2 neuronal precursor 937230 Homo sapiens
			. —					AA114075	cDNA clone 564098 3'
		H953251	8	15 7	22	84	Examples L76200	L76200	Homo sapiens guanylate kinase (GUK1) mRNA
Ē .	CAT GT GGAGT GGAGG	H955723		F.	37	4	Examples X00570	X00570	Human mRNA for precursor of apolipoprotein Cl
	CATGI GGCCCCAGGI	H962086		15 13	76	27	Examples L16510	L16510	Homo sapiens cathepsin B mRNA
2	168 CA1 61 6661 646CCA		.!	-				M14221	Human cathepsin B proteinase mRNA, complete cds
3		H975446	<u></u>	3	22	8	Examples L35240	L35240	Human enigma gene
	109 CAIGIGIGACCCCI	H976644	80	21 26	18	20	Examples L38941	138941	Homo sapiens ribosomal protein L34 (RPL34) mKNA
	TO CATCHGTGTGTTTGT	H978687	9	7 16	25	15	Examples X03473	X03473	Human gene for histone H1(0).
1						-	1	A A 034505	ZKZJĘUS.SI Soarcs pregnant uterus ivorus organicus septens conversiones
	1-2 CATSTATGGATCTC	H997944		=	11 711	7	Examples	Examples And to the	1/11/4

2131b06.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 723923		zk30c10.s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone	Т	yusouve, st nomin sapiens very contraction of the property	ESTU4595 Homo sapiens CDINA Clone nr DDA52	NTB1509 Normalized infant brain Bento Soares Homo sapiens cDNA	3'end similar to EST04595 H. sapiens cDNA clone HFBDX32	ze97h02 s1 Soares fetal heart NbHH19W Homo sapiens cDNA clone		2 Cosco 2 Cosco NAHTGBC Homo caniens cDNA clone 712204 3	П	ym05a09.s1 Homo sapiens cUNA clone 40073 3	H.sapiens mRNA for tyrosine kinase receptor.	Human mRNA for collagen VI alpha-l	H. sapiens gene for glutaminyl-tRNA synthetase	zk73h10,s1 Soares pregnant uterus NbHPU Homo sapiens cDNA clone		yz36b07.s1 Homo sapiens cDNA clone 285109 3'	l	П	H.sapiens (5) Ferriun H pseudogene.	Human mRNA for apoferritin H chain type	Human apoferritin H gene exons 2-4	Human ferritin heavy chain mRNA, complete cds	Human ferritin heavy chain mRNA, complete cds				2b17a08,s1 Homo sapiens cDNA clone 302294 3'	zt33d02.s1 Soares ovary tumor NbHOT Homo sapiens cDNA clone 724131			347396 3'
	AA235464	0000	AAU3/024	4297CH	T06706		T16635		AA02667	0000	AA280283	H10141	X66029	X15880	X72414		AA04456	N71899		AA400793	X80336	X00318	X03488	M97164	L20941	X02493	M11948	M17733	N78832		AA411095		W81693
				Examples H53629					Examples AA026678				Examples X66029	Examples X15880			Examples AA044568				Examples X80336					Examples X02493			Examples N78832				
				<u> </u>					v.				17	-			24	1			369					101							
		igspace		의				Ţ	3 24				8				16 10	l	-		84 235	L	_	igert	-	17 183	+	-	13			+	
-		┼		-				+	4	_	_	\vdash	6	L	+	+	=	1	╁╴		75 8	1	-	\dagger	+	1	1	╁	╁	+		\dagger	
 		\dagger	1	9				\dagger	٠,				te	†	+	+	4	†	†		202		T	T	1	901 86	1	T	6	1			
				H1003443					H1014660				H1021276	003120111	H1023220		H1024568	2011			H1026814					H1027595	2017		777770111	COLU	,	<u> </u>	
				CATGTTCATTGTAGA						21.416111161671				S STRECCCOOL				. TGTTGGAGAICIC				3 (A) 61 1 66 6 61 1 1 C					CATGIT GGI GAAGGA			IND CATGTTTCCCTCAAA			

Liman besingtone clathrin light-chain a mRNA	- 1	Human lymphocyte clathrin light-chain A mktwa	١	-	Human connective tissue growth factor mkNA	10,0001	wi78c08 s1 Homo sapiens cDNA clone 44273 3	İ	FST94173 Homo sapiens CDNA 3 end simulal to indire	1. OLI 170 A	A A 2 5 2 18 12 5 3 1 Soares NhHMPu SI Homo Sapiens CDIA CIONE CONTROL			
127001	Examples M20471	NA20472	- Carri	Examples X78947	0567111	07+10	106400	LICOLIZ	T25057	307061	L A A S C S A I	766744		
-	_	\dagger	-	_	\mid		-				-			
	1 6 3 7 I			0 0	1						t	_		
	3	+		0	1		+	_			+	_	1	
	2	+	_	٦	+	_	1				1		$\frac{1}{2}$	
	7078701L	111000111		11041504	10014010			H1044725	,				1	
		SI CATGTTTCCTTCCTT			INTICATGETT GCACCITIE				NATION 161 I MANA					

Table 5 - Transcripts increased in pancreas and colorectal cancer

SAGE tag that were elevated in both in coloreactal and pancreatic tumor, and are likely to be specific for tumor in general.

	Tag Semience		Tag Number Accession	Accession		rion
	TOATE TEGABATEAC	U	-950498 M10629	M10629	Human alpha-1 collagen	Ϋ́
1	TONTO CACTTORAGG	U	-294155	294155 042376	uman retinoic acid	Human retinoic acid induced RIG-E precursor (E) mR
	מוני בשנו בשנים			056145	uman thymic shared	Human thymic shared antigen-1/stem cell antigen-2
ſ	CATC ATCTCABGAG	T(A)	-243747	243747 J03040	Human SPARC/osteonectin mRNA,	tin mRNA, complete cds.
	2			M25746	Human osteonectin g	- 1
	CATG GCCCAAGGAC	U	-610466	610466 X53416	uman mRNA for acti	Human mRNA for actin-binding protein (filamin) (AB
7 4	CATC ATCTTGTTAC	L	-229106 X02761	X02761	uman mRNA for fibr	ctin (FN precursor).
				K00799	human fibronectin (fn)	3' coding region a
	ACATG GTGCGCTGAG	U	-760291 X58536	X58536	man mRNA for HLA	I locus C heavy
<u>`</u>				M26432	Human MHC class I HLA-C.1	gene, complete cds.
	CATG ACAGGCTACG	ß	-76231	-76231 M95787		nuscle protein (SM22) mkNA, com
				M83106	Human SM22 mRNA, 5'	end.
۵	a care greatertier	A	-769020 M77349	M77349	Human transforming	
0	POCATG GATTTCTCAG	U	-589267	589267 X53279	uman mRNA for plac	Human mRNA for placental-like alkaline phosphatase
				X55958	sapiens mRNA for	- 1
				J04948	Human alkaline phosphatase	(ALP-1) mRNA, comple
]	COTOTTACOA OTACIOT	E	-85882	85882 X57351	Human 1-8D gene fro	. 1
	באום שבבעוובובה			X02490	Human interferon-inducible	ducible mRNA (cDNA 1-8).
]	なしししかもししか しゅもし	C	-884181 X15804	X15804	Human mRNA for alph	alpha-actinin.
: :	TOTAL CTTCTGTGTA	, T	-515821	515821 080012	Human mRNA for KIAA	KIAA0190 protein.
	13 CATG ATGTABABA	F	-241665 M74090	M74090	Human TB2 gene mRNA,	31 end.
				J03801	Human lysozyme mRNA,	complete
				M19045	Human lysozyme mRNA,	complete cds.
7	CATG GGCAGAGGAC	U	-673954	673954 X17620	Human mRNA for Nm23	protein, involved in developme
				X75598	OH	
1	15 CATG AATATTGAGA	4	-53129	-53129 062962	Human Int-6 mRNA, c	
	16 CATG TTTTGATAA	A	-1048113 016891	016891	Human HepG2 3' region cDNA, clone	on CDNA, clone nmdzcii.
	17 CATG CAGCTGGCCA	Į.	-302741	-302741 X53743	H. sapiens mKNA for ilbuin-1	tabutan-i C.

			for Hib-nB antidens associated invarian
18 CATG GTTCACATTA	9	-774461 X00497	MKNA 101 nun-Dix antergene
		M13560	chain gene,
	E	-2056 Y00345	- 1
		-58533 M61831	S-adenosylhomocysteine hydrolase
20 CATG AAIGCAGGCA		M61832	
adama of ones	(-918273 X16934	hB23 gene for B23 nucleophosmin.
CATO TOWNS		M28699	is nucleolar pho
		M23613	A, complete cds.
		M26697	Human nucleolar protein (B23) mRNA, complete cds.
22 CATE TTATEGRATE	E	-998030 M24194	Human MHC protein homologous to chicken B complex
5 L47		-274492 D23661	ĔĬ
		111567	~
TLUTTLE BECCTTTGTT	9	-155632 083174	for collagen binding protein 2.
		-97078 X57352	I
SATS	A	-1000193M17886	phosphoprotein
		30506	mkna, complere
22 CATE CGACCCACG	U	-398663 M12529	lete cds.
		K00396	oprotein E (epsilon 2 and 3 alleles)
SOUTH CAGATETTE	T 5	-298495 X56998	
		66695X	Human UbA52 placental mRNA for ubiquitin-52 amino
505450550000000000000000000000000000000	U	-501287 X07491	hypome
5		M91670	
ATTORDE ATTORDE	A	-256497 L14272	Human prohibitin (PHB) gene, exons 1-7.
	1	885655	prohibitin (human, mRNA, 1043 nt].
ACACCECCEC CEAC IS	0	-765573 062435	V)
11 CA10 010010		068041	Human breast and ovarian cancer susceptibility pro
ACCOUNT TOTAL	T	-883029 M24398	Human parathymosin mRNA, complete cds.
33 CATG ACTGGGTCTA	T 4	-125661 X58965	
5		M36981	Human putative NDP kinase (nm23-H2S) mRNA, complet
		116785	eo i
SATADAGAA DAGATAG	4	-33331 002032	partial
34 CATO AMONONIO		037230	complete
		043701	

				!	0010	Clone 01) liver expressed protein mR
					113/33	1.12 mRNA. comple
35	SCATG	ACATCATCGA	(-79065	LU65U5	protein S2
36	36 CATG	CTGTTGGTGA	H	-507577 014530	14530	10g or yeast ilbosomar proton 17
	37 CATG		1	-249854 X57959	187959	for ribosomal process
	2				X57958	o
					X52967	L/.
			-	1	116558	nRNA, complete
;	1	I SERTITION	4	-6551151	L06498	0 (RPS20) mRNA,
2 3	2 6	0001111000		-672265 L19527	19527	انج
	39 CA16	GGCAAGAAGA	+		L25346	sapiens ribosomal protein L27 (homologue of
	1	CTCTTCGAGA	A	-490889 Y00433	(00433	
2	2	5.55.15.15			Y00483	- 1
	\perp				x13710	H. sapiens unspliced mRNA for glutathione peroxidas
			1	Î	x13709	Human gpx1 mRNA for gluthatione peroxidase.
			+		M21304	
			1	74540x 224702	78677	Human liver mRNA fragment DNA binding protein UPI
4	1 CATG	41 CATG CTGTTGATTG		CC \$ 100-	7 7 5 0 0 1 1	Human clone C4E 3.2 (CAC)n/(GTG)n repeat-containin
		١	+	10000	1757	H sapiens S19 ribosomal protein mRNA, complete cds
42	2 CATG	CTGGGTTAAT	A	1C110M 47/705-	107701	i
4	43 CATG	ATGCTGGTA	<u>-</u>	-239533 X1/200	007/17	HILLS TOND FOR EDSTEAD BALL VIEWS SMAll RNAS (EBER
4	44 CATG	GATGCTGCCA	4	-5835/3X5935-	10560	TOT UNIVERSE
	_				121756	complete cds.
	_				200/10	1
	_				5/6343	income interior of mRNA, complete cds.
4	45 CATG	CCTTCGAGAT	٥	-390692 014970	014970	complete cds.
4	46 CATG	CTCCTCACCT	9	-482584 016811	118910	in mRNA.
			1	20000	023/02	on factor
4	47 CATG	TGTGTTGAGA	5	C700/6-	V16872	Human DNA for elongation factor 1-alpha (clone lam
					x03558	mRNA for elongation factor
	_		\top		017182	HepG2 3' region MboI cDNA,
	-				017245	HepG2 3' region MboI cDNA,
	_				017259	3' region Mbol cDNA, clone
	-		T		017276	Human HepG2 3' region MboI cDNA, clone hmd6al2m3.

	130000	uman elemention factor 1 alpha mRNA, 3' end.
	M2 / 304	cloudation factor 1-alpha
	94C67W	TI-1 mRNA complete co
	L41490	Saptens oncogene its and a complete
	L41498	COMPLETE
A P CATG TTACCATATC A	-988366 U57846	Human ribosomal protein L39 mRNA, complete cds.
CATC GCTGCTGG	-621035 X71973	V) I
CTTGGAAAA	-383489 226876	
TACAAGAGGA	-803369 X69391	n L6.
	-803369 D17554	AXREB107,
	-803369 S71022	neoplasm-related C140 product [human, thyroid carc
SO CATE AACGACCICG I	-24951 V00598	
	-24951 V00599	oding beta-tubulin.
53 CATG CCCTGCCTTG T	-358783 X55110	moting proce
SA CATE CCAGGGGG A	-346761 038846	Human stimulator of TAR RNA binding (SKB) mKNA, CO
	016933	region cDNA, clone nmd
G CATCHOOM OF A G	-148949 211692	H.sapiens mRNA for elongation factor 4.
	-416261 X73974	
	053660	• 1
ET CATE CTABABABA A	-458753 M33680	TAPA-1 MKN
CATE CECTGATGTG	-686319 009510	synthetase mRNA, complete
200000000000000000000000000000000000000	009587	mRNA, complete co
	D30658	Human T-cell mRNA for glycyl tRNA synthetase, comp
SOUTH ATTENDED A	-253260 X55954	Human mRNA for HL23 ribosomal protein homologue.
	X52839	Human mRNA for ribosomal protein L17.
SOLCATE GAAAAATGGT T	-524524 X61156	H. sapiens mRNA for laminin-binding protein.
	X15005	
	043901	
	303799	g protein
	M14199	mRNA, 5' e
SICACTOACTO	-302367 087735	Human mRNA for ribosomal protein L14, complete cds
	L10376	clone CTG-B33) mRNA sequence.
	880520	beat-containing
G CATABATT G	-200576 014973	Human ribosomal protein S29 mRNA, complete cds.

			13	L31610	Homo sapiens (clone cori-1cl5) S29 ribosomai proce
1	SOLUTION DEAD	A 557	-55227 228407	Г	H. sapiens mRNA for ribosomal protein L8.
3	CAIG AAICCIG		-51925 M64716	Τ	Human ribosomal protein S25 mRNA, complete cds.
6.4	CATG AATAGGICCA		21,77	T	
		, (C,	8×11	-1 XB3412	H sapiens Bl mRNA for mucin.
65	CATG AAAAAAAA		23	Т	H. sapiens FRGAMMA mRNA (819bp) for folate receptor
			23	Ţ	H. sapiens FRGAMMA' mRNA for folate receptor (817bp
			1X	Γ	H. sapiens mRNA for lung amiloride sensitive Na+ ch
			00		Human FR-gamma' mRNA, complete cds.
			00	008471	Human folate receptor 3 mRNA, complete cds.
			U4	048697	Human mariner-like element-containing mRNA, clone
			D2	П	spha
			MS	M55914	Human c-myc binding protein (MBP-1) mRNA, complete
			10	Т	
			57	Τ	calmitine-mitochondrial calcium-binding protein (h
			72	T	transcript ch138 [human, RF1, RF48 stomach cancer c
			9×	Γ	H. sapiens mRNA for mitochondrial phosphate carrier
			-335945 X79238	Ī	H.sapiens mRNA for ribosomal protein L30.
9	CATG CCAGAACAGA	۱		Τ	Human thymidylate kinase (CDC8) mRNA, complete cds
		١	246831X80822	T	H sapiens mRNA for ORF.
67	CATG AAGGTGGAGG	1	73000 C00FF	Т	Human cyclophilin-related processed pseudogene.
8	68 CATG CCTAGCTGGA	I Was	A COCCIO	Τ	Human cyclophilin-related processed pseudogene.
			2 ×	Т	Human cyclophilin-related processed pseudogene.
			XSX	X52851	Human cyclophilin gene for cyclophilin (EC 5.2.1.8
			YO	Π	
18	COLOUR CARCACATO	A JUL	-528694 X63527	53527	119.
o	באום משכשכו		SS	S56985	
,	CATE BAGGAGATGG	7.GG G	-41531 X6	X69181	H.sapiens mRNA for ribosomal protein L31.
			[X]	X15940	Human mRNA for ribosomal protein L31.
-	applanded of a	A 400	-171113 229650	29650	
-	2		[D]	017233	ទ
		7	-177610 X08096	96080	Human GST pi gene for glutathione S-transferase pi
7,	72 CATG AGGICCIAGE	١			

			2000	uman mont for class Pi glutathione S-transferase
			X06547	MUMBIL MINNA LOL CLOSE AL CORD. TOT ANNUAL MUMBILLE
			X15480	Human mRNA for anionic glutathione-5-transferase (
			X08058	Human glutathione S-transferase pi gene.
			U12472	glutathione S-transferase (GST phi)
			021689	Human glutathione S-transferase-Plc gene, complete
			062589	Human glutathione S-transferase Plc (GSTplc) mRNA,
			M69113	. 1
			M24485	3ST-pi) glutatl
1	73 CATE TEGTETTERS 6	-965603	965603 X69150	protei
			M96153	
			L06432	٦١
٦	A CATCAACATCT C	-475448 M17885	M17885	tein PO m
-	TE CATE CITETABLES G	-769045 L25899	L25899	plete c
- -		-174037 X58125	X58125	Human (D9S55) DNA segment containing (TG)24 repeat
			D17268	Human HepG2 3' region MboI cDNA, clone hmd5h09m3.
			M73791	novel gene mRNA, comp
			M64241	s tumor-related protein (QM) mRNA,
			835960	3' region] (hum
	1 つつつままなつつ つまもつ	-671654	671654 M17887	Human acidic ribosomal phosphoprotein P2 mRNA, com
			M11147	Human ferritin L chain mRNA, complete cds.
			M12938	Human ferritin light subunit mRNA, partial cds.
			M10119	Human ferritin light subunit mRNA, complete cds.
	O See a Contraction of the Contr	-246019	246019 X04409	ng protein G(s) alpha-
			X04408	Human mRNA for coupling protein G(s) alpha subunit
			80095X	Human GSA mRNA for alpha subunit of GsGTP binding
			x07036	
			M21142	alpha
			M14631	Human guanine nucleotide-binding protein G-s, alph
	A ATSTOCATET A	-968173	3236832	H.sapiens (xs31) mRNA, 835bp.
			K00558	human alpha-tubulin mRNA, complete cds.
j	C SORCIOLOGIC OF ACTION	-955718	X56494	H.sapiens M gene for M1-type and M2-type pyruvate
c			M23725	Human M2-type pyruvate kinase mRNA, complete cds.
			M26252	Human TCB gene encoding cytosolic thyroid hormone-
			-	

			74774 XA7747	H. sapiens rpS8 dene for ribosomal protein S8.
£ 5	CATG TANTAAAGGI		-602315 X89401	for
	CATE GCATAATAGE		014967	Human ribosomal protein L21 mRNA, complete cds.
			025789	Human ribosomal protein L21 mRNA, complete cds.
			138826	Homo sapiens L21 ribosomal protein gene, partial c
Ę	CATG TACCATCAAT	SAAT A	-807748 X53778	- 1
			034995	Human normal keratinocyte substraction library mRN
			302642	Human glyceraldehyde 3-phosphate dehydrogenase mRN
			M36164	dehydrogenase
			M33197	dehydrogenase (
9.4	CATG ATTIGICCCA	SCCA G	-260949 X14957	
	2		X14958	Human hmgI mRNA for high mobility group protein Y.
			M23614	Human HMG-I protein isoform mRNA (HMGI gene), clon
			M23619	- 1
			117131	- HMG-
			M23615	Human HMG-Y protein isoform mRNA (HMGI gene), clon
			M23616	Human HMG-Y protein isoform mRNA (HMGI gene), clon
			M23617	Human HMG-Y protein isoform mRNA (HMGI gene), clon
			M23618	Human HMG-Y protein isoform mRNA (HMGI gene), clon
3	CATG GAGGGAGITT	STTT C	-567488 014968	~ !
3 2	CATG	1 266C T	-416106 012465	Human ribosomal protein L35 mRNA, complete cds.
2 6	CATG GTGAAACCCA	CCCA ALL	-753749 263072	~ 1
88	CATG GTGAAACCCA	CCCA ALL	-753749 X16294	
89	CATG AAGACAGTGG	sree c	-33979 X66699	- 1.
			T06499	Homo sapiens ribosomal protein L37a (RPL37A) mRNA,
			L22154	Human ribosomal protein L37a mRNA sequence.
96	CATG CCCCAGCCAG	ccag T	-348755 X55715	
			014990	MKNA,
			014991	orotein S3 (rpS3) mRNA, c
			014992	rpS3) mRNA,
			842658	
ē	CATG TGGGCAAAGC	AAGC C	-959498 X63526	
			211531	H.sapiens mRNA for elongation factor-1-gamma.

				-		THE STATE OF
					M55409	reference process minister
١	E & U	ATAGOODACT OFFI	4	-928269 M10036	M10036	somerase mknA, complete
75	2147	TOWORDS TO	: 0	-549145	9145 058682	complete cds.
93	3 CATG	GALGALACGA	,		M58458	S4 (RPS4X)
	1				M22146	
-			6	-26261	223063	y fact
94	CATC	AACGCGGCCA	-		110612	n glycosylation-inhi
					M95775	>
1	1				1,19686	migration inhibitory f
	\downarrow				M25639	migration inhibite
- [1		L	-935680 X03342	X03342	L32.
2	2 CA16	100000	,		K03002	Human mRNA from chromosome 15 gene with homology t
Ĭ	06 000	CACAAACGGT	A	-278636 U57847	US7847	Human ribosomal protein S27 mRNA, complete cds.
	2				L19739	1 (MFSI) (IIINA)
0	07 CATG	GGAGTGGACA	F	-667269	L11566	ans ribosomal protein bid (Nrbid) minor
٦١٥			S	-615043 254999	254999	island UNA genomic Misel Ilayment
^	2				257572	DNA genomic Msei Itagment
	<u> </u>				256073	genomic Msel Ilagineile,
	-				X53505	ribosomal protein 512.
P	100		Ĺ	-696375	M92381	
و	99 CATG	CATG GGGGAAA1CG	,			Human thymosin beta-10 mRNA, complete cds.
			,	500350	5003501114969	ribosomal protein L28 mRNA, compl
9	OCATG	s gcagccarcc	٥	20000	017257	
	-		-	07777X 15897-	07777X	6 mRNA.
2	CATG	CATG TARGGAGCIG			X69654	H. sapiens mRNA for ribosomal protein S26.
			4	-672342		cds.
2	102 CA16	GOCHAGOCCC			X79239	protein 513.
	-				L01124	Human ribosomal protein S13 (RPS13) mRNA, complete
- :			C	-775658 X65923	X65923	
2	103 CA16	0110001000	ì		U02523	ide repea
1	100	ยองสอบนอบบ	U	-374027	374027 M60854	plete cds.
-	104 CA16		,	-1027448 212962	212962	OSOMAL
	CATG	G TTGGTCCICI	,		564030	L41 ribosomal protein homolog (clone 786) (human,
	_					

- 1	200017 0012000	Himan mRNA for cytokeratin 18.
105 CATG CAAACCATCC A	-2634 /8 A12663	
	X12876	יייייייייייייייייייייייייייייייייייייי
	X12881	Human mRNA for cytokeratin 18.
	M24842	Human keratin 18 (K18) gene, complete cds.
	M26325	Human cytokeratin 18 mRNA, 3' end.
	M26326	Human keratin 18 mRNA, complete cds.
	M26327	cytokeratin 1
S TOOTTTOO DEAD SOL	-161624 X53777	L23 mRNA for putative riboso
	-177315 D86979	Human male bone marrow myeloblast mRNA for KIAA022
	X55923	Human DNA for Alu element P1N6.
	8496X	
	X12544	~ I
	686112	ne 6 HindIII fr
	011831	Human clone 2102V-I chromosome 18p telomeric seque
	1112580	Human Alu repeat sequence A3.
	000710	
	012382	numan at the company of
	012583	Alu repear seque
	014694	repeat,
	014695	Human Alu-Sb2 repeat, clone HALUSBI5.
	014696	Human Alu-Sb2 repeat, clone HALUSB27.
	1114697	Alu-Sb2
	1114698	
	1114699	
	111 4 700	Alu-Sb2
	1114701	repeat,
	1114704	
	1114706	Human Alu-Sb2 repeat, clohe HUM-10.
	014707	HUM-7.
	300120	Human (Lawn) c-myc proto-oncogene, complete coding
	134653	Homo sapiens platelet/endothelial cell adhesion mo
	M37521	Human XV2c gene.
	861789	ł
	\$73483	phosphorylase kinase catalytic subunit PHKG2 homol

	102512	cholinesterase (Alu element) (numan, institudi dat
	575337	(Y Alu polymorphism, YAP, polymorphic subfamily-3)
CATE GEOTTONE	-695980 249148	protein L
	010248	č V
	049083	Human cell surface heparin binding protein HIP mRN
	016992	
	D16911	e hmd3b09.
	J03537	
	M20020	Human ribosomal protein S6 mRNA, complete cds.
109 CATG ACGITCICIT C	-114144	EST
110 CATG TCTCCATACC C	-906438	EST
111 CATG GACTGCGTGC C	-555450	EST
112 CATG CTTAATCCTG A	-508767	EST
113 CATG GGTTGGCAGG G	-719435	EST
114 CATG GCCTCTGCC A	-613862	EST
115 CATG AACAGAAGCA A	-18469	LS3
116 CATG CTGCCGAGCT C	-497192	EST
117 CATG TTCCTCGGGC A	-1007018	EST
118 CATG AACTAATACT A	-28872	EST
119 CATG TAGATAATGG C	-822331	EST
120 CATG GCCACACCCC A,C	-607318	EST
121 CATG GAACCCTGGG A	-529899	EST
122 CATG AACTAAAAAA A	-28673	EST
123 CATG GAAATGTAAG A	-528067	EST
124 CATG ACTCCAAAAA A	-119809	EST
125 CATG GTTCGTGCCA A	-77109	EST
126 CATG TTACCTCCTT C	-989024	EST
127 CATG GCACAAGAAG A	-594051	
128 CATG CCCTGGGTTC T	-359102	EST
129 CATG GCCTGTATGA G	-621369	EST
30 CATG CCCGTCCGGA A	-355689	EST
131 CATG AGGAAAGCTG C	-163999	EST
132 CATG TCAGATCTTT G	-861056	EST

EST EST EST EST

			;	2
-618195	ပ	GCGTGTCCG	1 26 CATG	3,26
01010	₩.		135 CATG	135
769605				
-02/20-	ပ	TCACCCACAC	134 CATG	134
057363			2	133
122000	_	CCAGGAGGAA	123 CATG	133
190966				

Isolation of partial cDNA (3' fragment) by 3' directed PCR reaction

This procedure is a modification of the protocol described in Polyak et al. (1997) Nature 389:300. Briefly, the procedure uses SAGE tags in PCR reaction such that the resultant PCR product contains the SAGE tag of interest as well as additional cDNA, the length of which is defined by the position of the tag with respect to the 3' end of the cDNA. The cDNA product derived from such a transcript driven PCR reaction can be used for many applications.

RNA from a source believed to express the cDNA corresponding to a given tag is first converted to double-stranded cDNA using any standard cDNA protocol. Similar conditions used to generate cDNA for SAGE library construction can be employed except that a modified oligo-dT primer is used to dreive the first strand synthesis. For example, the oligonucleotide of composition 5'-B-TCC GGC GCG CCG TTT T CC CAG TCA CGA(30)-3', contains a poly-T stretch at the 3' end for hybridization and priming from poly-A tails, an M13 priming site for use in subsequent PCR steps, a 5' Biotin label (B) for capture to strepavidin-coated magnetic beads, and an AscI restriction endonuclease site for releasing the cDNA from the streptavidin-coated magnetic beads. Theoretically, any sufficiently-sized DNA region capable of hybridizing to a PCR primer can be used as well as any other 8 base pair recognizing endonuclease.

cDNA constructed utilizing this or similar modified oligo-dT primer is then processed exactly as described in U.S. Patent No. (insert) up until adapter ligation where only one adapter is ligated to the cDNA pool. After adapter ligation, the cDNA is released from the streptavidin-coated magnetic beads and is then used as a template for cDNA amplification.

Various PCR protocols can be employed using PCR priming sites within the 3' modified oligo-dT primer and the SAGE tag. The SAGE tag-derived PCR primer employed can be of varying length dictated by 5' extension of the tag into the adaptor sequence. cDNA products are now available for a variety of applications.

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This technique can be further modified by: (1) altering the length and/or content of the modified oligo-dT primer; (2) ligating adaptors other than that previously employed within the SAGE protocol; (3) performing PCR from template retained on the streptavidin-coated magnetic beads; and (4) priming first strand cDNA synthesis with non-oligo-dT based primers.

Isolation of cDNA using GeneTrapper or modified GeneTrapper Technology

The reagents and manufacturer's instructions for this technology are commercially available from Life Technologies, Inc., Gaithersburg, Maryland. Briefly, a complex population of single-stranded phagemid DNA containing directional cDNA inserts is enriched for the target sequence by hybridization in solution to a biotinylated oligonucleotide probe complementary to the target sequence. The hybrids are captured on streptavidin-coated paramagnetic beads. A magnet retrieves the paramagnetic beads from the solution, leaving nonhybridized single-stranded DNAs behind. Subsequently, the captured single-stranded DNA target is released from the biotinylated oligonucleotide. After release, the cDNA clone is further enriched by using a nonbiotinylated target oligonucleotide to specifically prime conversion of the single-stranded target to double-stranded DNA. Following transformation and plating, typically 20% to 100% of the colonies represent the cDNA clone of interest. To identify the desired cDNA clone, the colonies may be screened by colony hybridization using the 32P-labeled oligonucleotide as described above for solution hybridization, or alternatively by DNA sequencing and alignment of all sequences obtained from numerous clones to determine a consensus sequence.

The genes which are identified herein as being differentially expressed in normal and cancer cells can be used diagnostically and prognostically. Transcription levels in a test sample suspected of being neoplastic can be determined and compared to the levels in normal colon cells. The test sample may be from any tissue suspected of neoplasia, and particularly from either suspected colorectal or suspected pancreatic cancer cells. The control cells for

the purposes of comparison are normal cells, preferably of the same tissue type as the test sample, e.g., colon cells, or pancreatic duct epithelial cells. Upregulation of transcription or downregulation of transcription is therefore diagnostic of the neoplastic state, depending on what gene is used as a test reagent. Similarly, transcription levels can be monitored to assess patent responses to anti-tumor therapies. Transcription levels will also provide prognostic information. For example, the level of transcription in a test sample can be compared to levels found in bona fide normal and tumor cells. More extreme deviations from normal expression levels indicate a poorer prognosis.

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Transcription levels can be determined according to any means known in the art. These include, without limitation, Northern blots, nuclear run-on assays, in vitro transcription assays, primer extension assays, quantitative reverse transcriptase-polymerase chain reactions (RT-PCR), and hybrid filter binding assays. These techniques are well known in the art. See J.C. Alwine, D.J. Kemp, G.R. Stark, *Proc. Natl. Acad. Sci. U.S.A.* 74, 5350 (1977); K. Zinn, D. Di-Maio, T. Maniatis, *Cell* 34, 865 (1983); G. Veres, R.A. Gibbbs, S.E. Scherer, C.T. Caskey, *Science* 237, 415 (1987).

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Similarly, upregulated genes and downregulated genes can be detected by measuring expression of their protein products. This can be done by any means known in the art, including but not limited to Western (immuno) blot, enzyme linked immunoadsorbent assay, radioimmunoassay, and enzyme assay. Such techniques are well known in the art. Protein products can be detected in tissue samples of a test patient, using a suspect sample as a test sample, and a matched normal tissue sample from the same tissue type as a control. If normal tissue is not available then a closely related tissue type can be used. Desirably both the samples being compared will be from the same individual. Alternatively, aberrant expression levels of protein products can be detected in body samples, such as blood, serum, feces, urine, sputum. As a control, a normal matched sample can be used from a healthy individual. Aberrant expression levels of transcripts can also be detected in such body samples, particularly in blood and serum.

Probes for use in the assays for transcription levels of particular genes or sets of genes may be RNA or DNA. The probes will be isolated substantially free of other cellular RNAs or DNAs. If the reagent contains one probe then it will comprise at least 50% of the nucleic acids in the reagent composition. If the reagent contains more than one probe, then the proportion will decrease accordingly, so that specific probes will still comprise at least 50% of the nucleic acids in the reagent composition.

Probes can be labeled according to any means known in the art. These may include radioactive labels, fluorescent labels, enzymatic labels, and binding partner labels such as biotin. Means for labeling and detecting probes are well known in the art. Probes comprise at least 10, 11, 12, 15, 20, or 30 contiguous nucleotides of a selected gene.

This invention provides proteins or polypeptides expressed from the polynucleotides of this invention, which is intended to include wild-type and recombinantly produced polypeptides and proteins from procaryotic and eucaryotic host cells, as well as muteins, analogs and fragments thereof. In some embodiments, the term also includes antibodies and anti-idiotypic antibodies.

It is understood that functional equivalents or variants of the wild-type polypeptide or protein also are within the scope of this invention, for example, those having conservative amino acid substitutions. Other analogs include fusion proteins comprising a protein or polypeptide.

The proteins and polypeptides of this invention are obtainable by a number of processes well known to those of skill in the art, which include purification, chemical synthesis and recombinant methods. Full length proteins can be purified from a colon or pancreatic cell or tissue lysate by methods such as immunoprecipitation with antibody, and standard techniques such as gel filtration, ion-exchange, reversed-phase, and affinity chromatography using a fusion protein as shown herein. For such methodology, see for example Deutscher et al. (1999) Guide To Protein Purification: Methods In Enzymology (Vol. 182, Academic Press). Accordingly, this invention also

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provides the processes for obtaining these proteins and polypeptides as well as the products obtainable and obtained by these processes.

The proteins and polypeptides also can be obtained by chemical synthesis using a commercially available automated peptide synthesizer such as those manufactured by Perkin Elmer/Applied Biosystems, Inc., Model 430A or 431A, Foster City. The synthesized protein or polypeptide can be precipitated and further purified, for example by high performance liquid chromatography (HPLC). Accordingly, this invention also provides a process for chemically synthesizing the proteins of this invention by providing the sequence of the protein and reagents, such as amino acids and enzymes and linking together the amino acids in the proper orientation and linear sequence.

Alternatively, the proteins and polypeptides can be obtained by well-known recombinant methods as described, for example, in Sambrook et al., (1989), supra, using the host cell and vector systems described above.

Also provided by this application are the polypeptides and proteins described herein conjugated to a detectable agent for use in the diagnostic methods. For example, detectably labeled proteins and polypeptides can be bound to a column and used for the detection and purification of antibodies. They also are useful as immunogens for the production of antibodies as described below. The proteins and fragments of this invention are useful in an in vitro assay system to screen for agents or drugs, which modulate cellular processes.

The proteins of this invention also can be combined with various liquid phase carriers, such as sterile or aqueous solutions, pharmaceutically acceptable carriers, suspensions and emulsions. Examples of non-aqueous solvents include propyl ethylene glycol, polyethylene glycol and vegetable oils. When used to prepare antibodies, the carriers also can include an adjuvant that is useful to non-specifically augment a specific immune response. A skilled artisan can easily determine whether an adjuvant is required and select one. However, for the purpose of illustration only, suitable adjuvants include, but

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are not limited to Freund's Complete and Incomplete, mineral salts and polynucleotides.

This invention also provides a pharmaceutical composition comprising any of a protein, analog, mutein, polypeptide fragment, antibody, antibody fragment or anti-idiotipic antibody of this invention, alone or in combination with each other or other agents, and an acceptable carrier. These compositions are useful for various diagnostic and therapeutic methods.

Antibodies can be generated using the proteins encoded by the transcripts identified by the tags disclosed herein. Use of all or portions of the protein as immunogens is routine in the art. Similarly, fusion proteins can be used as immunogens. Antibodies can be affinity purified using the proteins or portions thereof used as immunogens. Similarly, monoclonal antibodies specifically immunoreactive with the protein sequences of the invention can be generated according to techniques which are well known in the art.

Antibodies can be used analytically to quantitate the expression of particular transcripts identified herein as upregulated or downregulated in cancer. In addition, antibodies can be conjugated or non-covalently linked to cytotoxic agents, such as cytotoxins, radionuclides, chemotherapeutic drugs, etc. Such antibodies can be used therapeutically to specifically target cancer cells in which the protein antigens are upregulated. These include the proteins encoded by the transcripts identified by the tags shown in Tables 2, 4, and 5. Means of making such linked cytotoxic antibodies and of administering the same are well known in the art.

Also provided by this invention is an antibody capable of specifically forming a complex with the proteins or polypeptides as described above. The term "antibody" includes polyclonal antibodies and monoclonal antibodies. The antibodies include, but are not limited to mouse, rat, and rabbit or human antibodies.

Laboratory methods for producing polyclonal antibodies and monoclonal antibodies, as well as deducing their corresponding nucleic acid sequences, are known in the art, see Harlow and Lane (1988) supra and

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Sambrook et al. (1989) supra. The monoclonal antibodies of this invention can be biologically produced by introducing protein or a fragment thereof into an animal, e.g., a mouse or a rabbit. The antibody producing cells in the animal are isolated and fused with myeloma cells or heteromyeloma cells to produce hybrid cells or hybridomas. Accordingly, the hybridoma cells producing the monoclonal antibodies of this invention also are provided.

Thus, using the protein or fragment thereof, and well known methods, one of skill in the art can produce and screen the hybridoma cells and antibodies of this invention for antibodies having the ability to bind the proteins or polypeptides.

If a monoclonal antibody being tested binds with the protein or polypeptide, then the antibody being tested and the antibodies provided by the hybridomas of this invention are equivalent. It also is possible to determine without undue experimentation, whether an antibody has the same specificity as the monoclonal antibody of this invention by determining whether the antibody being tested prevents a monoclonal antibody of this invention from binding the protein or polypeptide with which the monoclonal antibody is normally reactive. If the antibody being tested competes with the monoclonal antibody of the invention as shown by a decrease in binding by the monoclonal antibody of this invention, then it is likely that the two antibodies bind to the same or a closely related epitope. Alternatively, one can pre-incubate the monoclonal antibody of this invention with a protein with which it is normally reactive, and determine if the monoclonal antibody being tested is inhibited in its ability to bind the antigen. If the monoclonal antibody being tested is inhibited then, in all likelihood, it has the same, or a closely related, epitopic specificity as the monoclonal antibody of this invention.

The term "antibody" also is intended to include antibodies of all isotypes. Particular isotypes of a monoclonal antibody can be prepared either directly by selecting from the initial fusion, or prepared secondarily, from a parental hybridoma secreting a monoclonal antibody of different isotype by using the sib selection technique to isolate class switch variants using the

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procedure described in Steplewski et al. (1985) Proc. Natl. Acad. Sci. 82:8653 or Spira et al. (1984) J. Immunol. Methods 74:307.

This invention also provides biological active fragments of the polyclonal and monoclonal antibodies described above. These "antibody fragments" retain some ability to selectively bind with its antigen or immunogen. Such antibody fragments can include, but are not limited to:

- (1) Fab,
- (2) Fab',
- (3) F(ab')2,
- (4) Fv, and
- (5) SCA

A specific example of "a biologically active antibody fragment" is a CDR region of the antibody. Methods of making these fragments are known in the art, see for example, Harlow and Lane, (1988) supra.

The antibodies of this invention also can be modified to create chimeric antibodies and humanized antibodies (Oi, et al. (1986) BioTechniques 4(3):214). Chimeric antibodies are those in which the various domains of the antibodies' heavy and light chains are coded for by DNA from more than one species.

The isolation of other hybridomas secreting monoclonal antibodies with the specificity of the monoclonal antibodies of the invention can also be accomplished by one of ordinary skill in the art by producing anti-idiotypic antibodies (Herlyn, et al. (1986) Science 232:100). An anti-idiotypic antibody is an antibody which recognizes unique determinants present on the monoclonal antibody produced by the hybridoma of interest.

Idiotypic identity between monoclonal antibodies of two hybridomas demonstrates that the two monoclonal antibodies are the same with respect to their recognition of the same epitopic determinant. Thus, by using antibodies to the epitopic determinants on a monoclonal antibody it is possible to identify other hybridomas expressing monoclonal antibodies of the same epitopic specificity.

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It is also possible to use the anti-idiotype technology to produce monoclonal antibodies which mimic an epitope. For example, an anti-idiotypic monoclonal antibody made to a first monoclonal antibody will have a binding domain in the hypervariable region which is the mirror image of the epitope bound by the first monoclonal antibody. Thus, in this instance, the anti-idiotypic monoclonal antibody could be used for immunization for production of these antibodies.

As used in this invention, the term "epitope" is meant to include any determinant having specific affinity for the monoclonal antibodies of the invention. Epitopic determinants usually consist of chemically active surface groupings of molecules such as amino acids or sugar side chains and usually have specific three dimensional structural characteristics, as well as specific charge characteristics.

The antibodies of this invention can be linked to a detectable agent or label. There are many different labels and methods of labeling known to those of ordinary skill in the art.

The antibody-label complex is useful to detect the protein or fragments in a sample, using standard immunochemical techniques such as immunohistochemistry as described by Harlow and Lane (1988) supra. Competitive and non-competitive immunoassays in either a direct or indirect format are examples of such assays, e.g., enzyme linked immunoassay (ELISA) radioimmunoassay (RIA) and the sandwich (immunometric) assay. Those of skill in the art will know, or can readily discern, other immunoassay formats without undue experimentation.

The coupling of antibodies to low molecular weight haptens can increase the sensitivity of the assay. The haptens can then be specifically detected by means of a second reaction. For example, it is common to use haptens such as biotin, which reacts avidin, or dinitropherryl, pyridoxal, and fluorescein, which can react with specific anti-hapten antibodies. See Harlow and Lane (1988) supra.

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The monoclonal antibodies of the invention also can be bound to many different carriers. Thus, this invention also provides compositions containing the antibodies and another substance, active or inert. Examples of well-known carriers include glass, polystyrene, polypropylene, polyethylene, dextran, nylon, amylases, natural and modified celluloses, polyacrylamides, agaroses and magnetite. The nature of the carrier can be either soluble or insoluble for purposes of the invention. Those skilled in the art will know of other suitable carriers for binding monoclonal antibodies, or will be able to ascertain such, using routine experimentation.

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Compositions containing the antibodies, fragments thereof or cell lines which produce the antibodies, are encompassed by this invention. When these compositions are to be used pharmaceutically, they are combined with a pharmaceutically acceptable carrier.

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The present invention also provides a screen for various agents which modulate the expression of a gene in a pancreatic or colon cell. To practice the method in vitro, suitable cell cultures or tissue cultures are first provided. The cell can be a cultured cell or a genetically modified cell in which a trancript from SEQ ID NOS:1-732, or their complements, is expressed. Alternatively, the cells can be from a tissue biopsy. The cells are cultured under conditions (temperature, growth or culture medium and gas (CO₂)) and for an appropriate amount of time to attain exponential proliferation without density dependent constraints. It also is desirable to maintain an additional separate cell culture; one which does not receive the agent being tested as a control.

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As is apparent to one of skill in the art, suitable cells may be cultured in microtiter plates and several agents may be assayed at the same time by noting genotypic changes, phenotypic changes or cell death.

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When the agent is a composition other than a DNA or RNA, the agent may be directly added to the cell culture or added to culture medium for addition. As is apparent to those skilled in the art, an "effective" amount must be added which can be empirically determined. When the agent is a polynucleotide, it may be directly added by use of a gene gun or

electroporation. Alternatively, it may be inserted into the cell using a gene delivery vehicle or vector as described above.

An agent is a potential therapeutic if it alters the expression of gene in the cell. Altered expression can be detected by assaying for altered mRNA expression or protein expression using the probes, primers and antibodies as described herein.

For the purposes of this invention, an "agent" is intended to include, but not be limited to a biological or chemical compound such as a simple or complex organic or inorganic molecule, a peptide, a protein (e.g. antibody) or a polynucleotide (e.g. anti-sense). A vast array of compounds can be synthesized, for example polymers, such as polypeptides and polynucleotides, and synthetic organic compounds based on various core structures, and these are also included in the term "agent". In addition, various natural sources can provide compounds for screening, such as plant or animal extracts, and the like. It should be understood, although not always explicitly stated that the agent is used alone or in combination with another agent, having the same or different biological activity as the agents identified by the inventive screen. The agents and methods also are intended to be combined with other therapies.

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The above disclosure generally describes the present invention. A more complete understanding can be obtained by reference to the following specific examples which are provided herein for purposes of illustration only, and are not intended to limit the scope of the invention.

EXAMPLE 1

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This example demonstrates the characterization of the general transcription of human colorectal epithelium, colorectal cancers, and pancreatic cancers.

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We used the recently developed SAGE (serial analysis of gene expression) method to identify and quantify a total of 303,706 transcripts derived from human colorectal (CR) epithelium, CR cancers or pancreatic cancers (Table 1A) (3). These transcripts represented approximately 48,741

different genes (4) that ranged in average expression from 1 copy per cell to as many as 5,300 copies per cell (5). The number of different transcripts observed in each cell population varied from 14,247 to 20,471. The bulk of the mRNA mass (75%) consisted of transcripts expressed at more than five copies per cell on average (Table 1B). In contrast, the majority (86%) of transcripts were expressed at less than 5 copies per cell, but in aggregate this low abundance class represented only 25% of the mRNA mass. This distribution was consistently observed among the different samples analyzed and was consistent with previous studies of RNA abundance classes based on RNA-DNA reassociation kinetics (Rot curves). Monte Carlo simulations revealed that our analyses had a 92% probability of detecting a transcript expressed at an average of three copies per cell (7).

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Table 1 - Summary of SAGE Analysis

A. Overall Summary

The second secon					The second name of the second na	The second second
	Normal	Colon	Colon	Pancreatic	Pancreatic	
	Colon	Tumors	Cell Lines	Tumors	Cell Lines	Total
Total Tags	62,168	80,878	60,373	61,592	58,695	303,706
Unique Genes ¹ 14,721 GenBank ² 8,753 (14,721 8,753 (59)	19,690 10,490 (53)	17,092 10,193 (60)	20,471 11,547 (56)	14,247 8,922 (63)	48,741 26,339 (54)

¹ Indicates the number of different genes represented by the total tags analyzed (4).

² Indicates the number of genes that matched an entry in GenBank. The number in parentheses indicates the corresponding percentage of total unique tags.

Table 1 - Summary of SAGE Analysis

B. Summarized by Abundance Classes*

	Normal	Colon	Colon	Pancreatic	Pancreatic Cell	ell
Copies/Cell	Colon	Tumors	Cell Lines	Tumors	Lines	Total
> 500	3	í		<u> </u>	;	
Unique Genes	62 (29)	54 (25)	54 (19)	32 (11)	70 (26)	25 (19)
GenBank	(56) 65	52 (96)	53 (98)	32 (100)	70 (100)	54 (98)
> 50 and < 500						
Unique Genes	645 (28)	470 (21)	618 (27)	657 (29)	585 (27)	595 (26)
GenBank	545 (84)	429 (91)	579 (94)	(60) (03)	529 (90)	553 (93)
> 5 and < 50						
Unique Genes	4,569 (27)	5,011 (29)	5,733 (34)	6,146 (36)	4,895 (31)	6,209 (30)
GenBank	2,893 (63)	3,204 (64)	3,682 (64)	4,054 (66)	3,168 (65)	4.241 (68)

	41,882 (25)	21,491 (51)	
	8,697 (16)	5,155 (59)	
	13,636 (24)	6,852 (50)	
	10,687 (20)	5,879 (55)	
	14,155 (25)	6,805 (48)	
	9,445 (16)	5,256 (56)	
o /I	Unique Genes	GenBank	

*For unique genes, the first number denotes the number of different genes (4) represented in the indicated abundance class. The number in parentheses indicates the mass fraction (X100) of total transcripts represented by the indicated abundance class. For GenBank entries, the first number indicates the number of different genes that matched an entry in GenBank in the indicated abundance class. The number in parentheses indicates the corresponding percentage of total genes.

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Many of the SAGE tags appeared to represent previously undescribed transcripts, as only 54% of the tags matched entries in GenBank (Table 1). Twenty percent of these matching transcripts corresponded to characterized mRNA sequence entries in GenBank, whereas 80% matched uncharacterized EST entries. As expected, the likelihood of a tag being present in the databases was related to abundance; GenBank matches were identified for 98% of the transcripts expressed at more than 500 copies per cell but for only 51% of the transcripts expressed at ≤ 5 copies per cell. Because the SAGE data provide a quantitative assay of transcript abundance, unaffected by differences in cloning or PCR efficiency, these data provide an independent and relatively unbiased estimate of the current completeness of publicly available EST databases.

EXAMPLE 2

This example demonstrates a comparison of the expression pattern of normal colon epithelium and primary colon cancers.

Comparison of expression patterns between normal colon epithelium and primary colon cancers revealed that the majority of transcripts were expressed at similar levels (Fig. 1A). However, the expression profiles also revealed 289 transcripts that were expressed at significantly different levels [P < 0.01, (8)]. Of these 289, 181 were decreased in colon tumors compared to normal colon (average decrease 10-fold; Fig. 1B; examples in Fig. 2A). Conversely, 108 transcripts were expressed at higher levels in the colon cancers than in normal colon (average increase 13-fold; Fig. 1C; examples in Fig. 2A). Monte Carlo simulations indicated that the analysis would have detected over 95% of those transcripts expressed at a 6-fold or greater level in normal vs. tumor cells or vice versa (9). Because relatively stringent criteria were used for defining differences [P < 0.01, (8)], the number of differences reported above is likely to be an underestimate.

EXAMPLE 3

This example demonstrates the similarities and differences between cancer cell line transcription and transcription of primary cancer tissues. To determine how many of the 289 differences were independent of the cellular microenvironment of cancers in vivo, SAGE data from CR cancer cell lines was compared to that from primary CR cancer tissues (Fig. 1B, 1C). Perhaps surprisingly, the majority of transcripts (130 of 181) that were expressed at reduced levels in cancer cells in vivo were also expressed at significantly lower levels in the cell lines (Fig. 1B). Likewise, a significant fraction of the transcripts expressed at increased levels in primary cancers were also expressed at higher levels in the CR cancer cell lines (Fig. 1C). Thus, many of the gene expression differences that distinguish normal from tumor cells in vivo persist during in vitro growth. However, despite these similarities there were also many differences. For example, only 47 of 228 genes expressed at higher levels in CR cancer cell lines were also expressed at high levels in the primary CR cancers.

In combination, comparing the expression pattern of CR cancer cells (in vivo or in vitro) to normal colon revealed 548 differentially expressed transcripts (Fig. 1B,C, Tables 2 and 3). The average difference in expression for these transcripts was 15 fold. Although the ability to detect differences is influenced by the magnitude of the variance with the power to detect smaller differences being less, 92 transcripts that were less than three fold different were identified among the 548 transcripts. However, those genes exhibiting the greatest differences in expression are likely to be the most biologically important.

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EXAMPLE 4

This example demonstrates the similarities and differences between colorectal cancer transcription and pancreatic cancer transcription.

To determine whether the changes noted in CR cancers were neoplasia or cell type specific, we performed SAGE on mRNA derived from pancreatic cancers. A total of 404 transcripts were expressed at higher levels in pancreatic cancers compared to normal colon epithelium (examples in Fig. 2B). The majority (268) of these transcripts were pancreas-specific (10) (Example in Fig. 2C) although 136 were also expressed at high levels in CR cancers. These 136 transcripts constituted 47% of the 289 transcripts increased in CR cancers relative to normal colon and are likely to be related to the neoplastic process rather than to the specific cell type of origin.

EXAMPLE 5

This example demonstrates the reproducibility of the transcription patterns observed among a larger number of cancer samples.

One question that arose from these data is the potential heterogeneity of expression between individual tumors. The SAGE data were acquired from two examples of each tissue type (normal colon, primary CR cancer, CR cancer cell line, etc.). To examine the generality of these expression profiles, we arbitrarily selected 27 differentially expressed transcripts and evaluated them in six to twelve samples of normal colon and primary cancers by Northern blot analysis (11). In general, expression patterns were very reproducible among different samples. Of 10 genes with elevated expression in normal colon relative to CR cancers as determined by SAGE, each was detected in the normal colon samples and was expressed at considerably lower levels in tumors (examples in Fig. 2A). Similarly, most of the genes identified by SAGE as increased in CR or pancreatic cancers were confirmed to be reproducibly expressed in the majority of primary cancers examined by Northern blot (examples in Fig. 2A). It is important to note, however, that there were differences among the cancers, with a few cancers exhibiting particularly high or low levels of individual transcripts. Such differences in gene expression

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undoubtedly contribute to the observed heterogeneity in biological properties of cancers derived from the same organ.

EXAMPLE 6

This example demonstrates the identities of some of the transcripts which were found to be differentially expressed in tumor and normal tissues. What are the identities of the differentially expressed genes? Of the 548 differentially expressed transcripts, 337 were tentatively identified through database comparisons. When tested, the great majority (93%) of these identifications proved to be legitimate (13), as expected from previous SAGE analyses. Although a large number of differentially expressed genes were identified, some simple patterns did emerge. For example, genes that were expressed at higher levels in normal colon epithelium than in CR tumors were often differentiation-related. These genes included liver fatty acid binding protein, cytokeratin 20, carbonic anhydrase, guanylin and uroguanylin, which are known to be important for the normal physiology or architecture of the colon epithelium (Table 2). On the other hand, genes that were increased in CR cancers were often related to the robust growth characteristics that these cells exhibit. For example, gene products associated with protein synthesis, including 48 ribosomal proteins, five elongation factors, and five genes involved in glycolysis were observed to be elevated in both CR and pancreatic cancers compared to normal colon cells. Although the majority of the transcripts could not have been predicted to be differentially expressed in cancers, several have previously been shown to be dysregulated in neoplastic The latter included IGFII, B23 nucleophosmin, the Pi form of glutathione S-transferase, and several ribosomal proteins which were all increased in cancer cells as previously reported. Likewise, Dra and gelsolin were both decreased in cancer as previously reported. Surprisingly, two widely studied oncogenes, c-fos and c-erbb3, were expressed at much higher levels in normal colon epithelium than CR cancers, in contrast to their up-regulation in transformed cells.

In summary, these data provide basic information necessary for understanding the gene expression differences that underlie cancer phenotypes. They additionally provide a necessary framework for interpreting the significance of individual differentially expressed genes. Although this study demonstrated that a large number of such differences exist (approximately 500 at the depth of analysis employed), it was equally remarkable that the fraction of transcripts exhibiting significant differences was relatively small, representing 1.5 % of the transcripts detected in any given cell type (26). The fact that many, but not all, of the differences were preserved during in vitro culture demonstrates the utility of cultured lines for examination of some aspects of gene expression, but also provides a note of caution in relying on such lines to perfectly mimic tumors in their natural environment. Finally, the finding that hundreds of specific genes are expressed at different levels in CR cancers, and that some of these are also expressed differentially in pancreatic cancers, provides a wealth of new reagents for future biologic and diagnostic experimentation.

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REFERENCES AND NOTES

- M. D. Adams, et al., Nature 377, supp. 28, 3 (1995); M. Schena, D. Shalon, R. W. Davis, P. O. Brown, Science 270, 467 (1995); J. Derisi, et al., Nature Genetics 14, 457 (1996); T. M. Gress, et al., Oncogene 13, 1819 (1996); D. J. Lockhart, et al., Nature Biotechnology 14, 1675 (1996); M. Schena, et al., Proc Natl Acad Sci USA 93, 10614 (1996).
- 2. V. E. Velculescu, L. Zhang, B. Vogelstein, K. W. Kinzler, Science 270, 484 (1995); V. E. Velculescu, et al., Cell 88, 243 (1997).
- of tags (30,000) were derived from two different patients for each tissue. For primary tumors (two CR carcinomas and two pancreatic adenocarcinomas), RNA was isolated from portions of tumors judged to contain 60%-90% tumor cells by histopathology. The cells grown in vitro were derived from CR (SW837, Caco2) and pancreatic (ASPC-1, PL45) cancer cell lines. CR epithelial cells were isolated from sections of normal colon mucosa from two patients using EDTA as previously described [S. Nakamura, I. Kino, S. Baba, Gut 34, 1240 (1993)]. Histopathology confirmed that the isolated cells were greater than 90% epithelial. Isolation of Poly-A RNA and SAGE was performed as previously described (2). SAGE data was analyzed by means of SAGE software and GenBank Release 95 as previously described (2).
- 4. A total of 69,393 different SAGE tags were identified among the 303,706 tags analyzed. A small fraction of these different tags were likely due to sequencing errors. SAGE analysis of yeast (2), wherein the entire genomic sequence is known, demonstrated a sequencing error rate of ~ 0.7%, translating to a SAGE tag error rate of 6.8% (1 0.993¹⁰). Because these sequencing mistakes are essentially random, they do not substantially affect the analysis although they could artificially inflate the number of unique genes identified. Therefore, to be conservative, we reduced our estimate of unique genes identified by this maximum tag error rate (e.g., 6.8% of 303,706 total tags). The number of different tags derived from the same gene due to alternative splicing was assumed to be negligible.

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- 5. Abundances can be simply determined by dividing the observed number of tags for a given transcript by the total number of tags obtained. An estimate of approximately 300,000 transcripts per cell was used to convert the abundances to copies per cell [N. D. Hastie, J. O. Bishop, *Cell* 9, 761 (1976)].
- 6. J. O. Bishop, J. G. Morton, M. Rosbash, M. Richardson, *Nature* **250**, 199 (1974); B. Lewin, Gene Expression Vol 2 (John Wiley and sons, New York 1980).
- 7. Computer simulations indicated that analysis of 300,000 tags would yield a 92 % chance of detecting a tag for a transcript whose expression was at least three copies per cell on average among the tissues examined and assuming 300,000 transcripts per cell.
- 8. To minimize the number of assumptions and to account for the large number of comparisons being made, Monte Carlo analysis was used for determining statistical significance. The null hypothesis was that the level. kind, and distribution of transcripts were the same for cancer and normal cells. For each transcript, 100,000 simulations were performed to determine the relative likelihood due to chance alone ("p-chance") of obtaining a difference in expression equal to or greater than the observed difference, given the null hypothesis. This likelihood was converted to an absolute probability value by simulating 40 experiments in which a representative number of transcripts (27,993 transcripts in each experiment) was identified and compared. The distribution of transcripts used for these simulations was derived from the average level of expression observed in the original samples. The distribution of the p-chance scores obtained in the 40 simulated experiments (false positives) was then compared to those obtained experimentally. Based on this comparison, a maximum value of 0.0005 was chosen for p-chance. This yielded a false positive rate that was no higher than 0.01 for the least significant p-chance value below the cutoff.
- 9. Two hundred simulations assuming an abundance of 0.0001 in one sample and 0.0006 in a second sample revealed a significant difference (P < 0.01, [8]) 95% of the time.

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- 10. It is not possible to obtain pancreatic ductal epithelium, from which pancreatic carcinomas arise, in sufficient quantities to perform SAGE. It is therefore not possible to determine whether these transcripts were derived from genes that were highly expressed only in pancreatic cancers or were also expressed in pancreatic duct cells.
- 11. Total RNA isolation and Northern blot analysis was performed as described [W. S. el-Deiry, et al., Cell 75, 817 (1993)].
- 12. A. H. Owens, D. S. Coffey, S. B. Baylin, Eds., Tumor Cell Heterogeneity: Origins and Implications (Academic Press, New York, 1982).
- Northern blot analyses were done on 45 of the 337 differentially expressed transcripts with tentative database matches. In three cases, the pattern of expression was not differentially expressed as predicted by SAGE and, for the purposes of this calculation, were presumed to represent incorrect database matches.
- 14. D. C. Rubin, D. E. Ong, J. I. Gordon, *Proc Natl Acad Sci U S A* 86, 1278 (1989); K. Okubo, J. Yoshii, H. Yokouchi, M. Kameyama, K. Matsubara, *DNA Res* 1, 37 (1994).
 - 15. R. Moll, et al., Differentiation 53, 75 (1993).
- 16. J. Sowden, S. Leigh, I. Talbot, J. Delhanty, Y. Edwards, Differentiation 53, 67 (1993).
- 17. F. J. de Sauvage, et al., Proc Natl Acad Sci USA 89, 9089 (1992).
 - 18. R. C. Wiegand, et al., FEBS Lett 311, 150 (1992).
- J. V. Tricoli, et al., Cancer Res 46, 6169 (1986); S. Lambert,
 J. Vivario, J. Boniver, R. Gol-Winkler, Int J Cancer 46, 405 (1990).
 - 20. W. Y. Chan, et al., Biochemistry 28, 1033 (1989).
- J. D. Hayes, D. J. Pulford, Crit Rev Biochem Mol Biol 30, 445
 (1995).
- G. F. Barnard, et al., Cancer Res 52, 3067 (1992); P. J. Chiao,
 D. M. Shin, P. G. Sacks, W. K. Hong, M. A. Tainsky, Mol Carcinog 5, 219 (1992); N. Kondoh, C. W. Schweinfest, K. W. Henderson, T. S. Papas,

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Cancer Res 52, 791 (1992); G. F. Barnard, et al., Cancer Res 53, 4048 (1993); M. G. Denis, et al., Int J Cancer 55, 275 (1993); J. M. Frigerio, et al., Hum Mol Genet 4, 37 (1995).

- 23. C. W. Schweinfest, K. W. Henderson, S. Suster, N. Kondoh, T. S. Papas, *Proc Natl Acad Sci U S A* 90, 4166 (1993).
- M. Tanaka, et al., Cancer Res 55, 3228 (1995); D. Medina, F.
 S. Kittrell, C. J. Oborn, M. Schwartz, Cancer Res 53, 668 (1993).
- A. D. Miller, T. Curran, I. M. Verma, Cell 36, 51 (1984); M.
 H. Kraus, W. Issing, T. Miki, N. C. Popescu, S. A. Aaronson, Proc Natl Acad Sci USA 86, 9193 (1989).
- 26. In the case of normal and neoplastic colon cancer tissue, 548 differentially transcripts were identified among the 36,125 unique transcripts.
 - 27. All references cited are hereby incorporated by reference herein.
- 28. Sequences tags in Tables 2-4 are consecutively numbered to form SEQ ID NOS: 1-732.

CLAIMS

1. A method of diagnosing colon cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 3;

identifying the first sample as neoplastic when the level of the at least one transcript is found to belower in the first sample than in the second sample.

2. A method of diagnosing colon cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2;

identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

- 3. The method of claim 1 wherein a comparison of at least two of said transcripts is performed.
- 4. The method of claim 2 wherein a comparison of at least two of said transcripts is performed.

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- 5. The method of claim 1 wherein a comparison of at least five of said transcripts is performed.
- 6. The method of claim 2 wherein a comparison of at least five of said transcripts is performed.
- The method of claim 1 wherein a comparison of at least ten of said transcripts is performed.
 - 8. The method of claim 2 wherein a comparison of at least ten of said transcripts is performed.
 - 9. The method of claim 1 wherein a comparison of at least twenty of said transcripts is performed.
 - 10. The method of claim 2 wherein a comparison of at least twenty of said transcripts is performed.
 - 11. The method of claim 1 wherein a comparison of at least thirty of said transcripts is performed.
- 15 12. The method of claim 2 wherein a comparison of at least thirty of said transcripts is performed.
 - 13. An isolated and purified human nucleic acid molecule which comprises a SAGE tag selected from SEQ ID NO:1-732.
 - 14. The nucleic acid molecule of claim 13 which is a cDNA molecule.

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- 15. The nucleic acid molecule of claim 13 wherein the SAGE tag is located at the 3' end of the molecule, adjacent to the 3'-most NlaIII restriction enzyme site.
- 16. An isolated nucleotide probe comprising at least 10 nucleotides of a human nucleic acid molecule, wherein the human nucleic acid molecule comprises a SAGE tag selected from SEQ ID NO: 1-732.
- 17. The probe of claim 16 which comprises the selected SAGE tag.
- 18. A diagnostic reagent for evaluating neoplasia of a colorectal tissue, comprising at least 2 probes according to claim 16.
- 19. The diagnostic reagent of claim 18 which comprises at least 5 probes according to claim 16.
 - 20. The diagnostic reagent of claim 18 which comprises at least 10 probes according to claim 16.
 - 21. The diagnostic reagent of claim 18 which comprises at least 20 probes according to claim 16.
 - The diagnostic reagent of claim 18 which comprises at least 30 probes according to claim 16.
 - 23. A diagnostic reagent for evaluating neoplasia of a colorectal tissue, comprising at least 2 probes according to claim 17.
 - 24. A method of diagnosing pancreatic cancer in a sample suspected of being neoplastic, comprising the steps of:

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comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a pancreatic tissue suspected of being neoplastic and the second sample is of a normal human colon tissue, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4;

identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

25. A method of diagnosing cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a tissue suspected of being neoplastic and the second sample is of a normal human tissue of the same tissue type, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 5;

identifying the first sample as neoplastic when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

26. A method to aid in the determination of a prognosis for a colon cancer patient, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic colonic tissue and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 3;

determining a poorer prognosis if the level of the at least one transcript is found to be lower in the first sample than in the second sample.

27. A method to aid in determining a prognosis for a patient with colon cancer, comprising the steps of:

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comparing the level of at least one transcript in a first tissue sample to a second sample, wherein the first sample is of a colonic cancer tissue and the second sample is of a normal human colonic tissue, and wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2;

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determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

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28. A method of diagnosing colon cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of expression of at least one protein in a

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comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 3;

identifying the first sample as neoplastic when the level of expression of the protein is found to be lower in the first sample than in the second sample.

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29. A method of diagnosing colon cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a colonic tissue suspected of being neoplastic and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 2;

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identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

30. A method to aid in determining a prognosis of a patient having pancreatic cancer, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic pancreatic tissue and the second sample is of a normal human colon tissue, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

31. A method to aid in providing a prognosis for a cancer patient, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic tissue and the second sample is of a normal human tissue of the same tissue type, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 5;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

32. A method of diagnosing pancreatic cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of expression of at least one protein encoded by a transcript in a first sample of a tissue to a second sample, wherein the first sample is of a pancreatic tissue suspected of being neoplastic and the second sample is of a normal human colon tissue, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4;

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

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33. A method of diagnosing cancer in a sample suspected of being neoplastic, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a tissue suspected of being neoplastic and the second sample is of a normal human tissue, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 5;

identifying the first sample as neoplastic when expression of the protein is found to be higher in the first sample than in the second sample.

34. A method to aid in the determination of a prognosis for a colon cancer patient, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic colonic tissue and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 3;

determining a poorer prognosis if the level of expression is found to be lower in the first sample than in the second sample.

35. A method to aid in determining a prognosis for a patient with colon cancer, comprising the steps of:

comparing the level of expression of at least one protein in a first tissue sample to a second sample, wherein the first sample is of a colonic cancer tissue and the second sample is of a normal human colonic tissue, and wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 2;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

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36. A method to aid in determining a prognosis of a patient having pancreatic cancer, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic pancreatic tissue and the second sample is of a normal human colon tissue, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

37. A method to aid in providing a prognosis for a cancer patient, comprising the steps of:

comparing the level of expression of at least one protein in a first sample of a tissue to a second sample, wherein the first sample is of a neoplastic tissue and the second sample is of a normal human tissue of the same tissue type, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 5;

determining a poorer prognosis if the level of expression is found to be higher in the first sample than in the second sample.

38. A method of treating a cancer cell, comprising the step of:

administering to a cancer cell an antibody which specifically binds to a protein encoded by a transcript identified by a tag selected from the group consisting of those shown in Tables 2, 4, and 5, wherein the antibody is linked to a cytotoxic agent.

39. An antibody linked to a cytotoxic agent, wherein the antibody specifically binds to a protein encoded by a transcript identified by a tag selected from the group consisting of those shown in Tables 2, 4, and 5.

40. A method of detecting colon cancer in a patient, comprising the steps of:

comparing the level of at least one protein in a first body sample to a second body sample, wherein the first sample is a body sample of the patient and the second sample is of a normal human, wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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identifying neoplasia when the level of the at least one protein is found to be higher in the first sample than in the second sample.

41. A method of detecting pancreatic cancer in a patient, comprising the steps of:

comparing the level of at least one protein encoded by a transcript in a first sample of a tissue to a second sample, wherein the first sample is of the patient and the second sample is of a normal human, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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identifying neoplasia when the level of the at least one protein is found to be higher in the first sample than in the second sample.

42. A method of detecting cancer in a patient, comprising the steps of:

comparing the level of at least one protein in a first sample to a second sample, wherein the first sample is of patient and the second sample is of a normal human, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 5, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

identifying neoplasia when the level of the at least one protein is found to be higher in the first sample than in the second sample.

43. A method to aid in determining a prognosis for a patient with colon cancer, comprising the steps of:

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comparing the level of at least one protein in a first sample to a second sample, wherein the first sample is of a colonic cancer patient and the second sample is of a normal human, wherein the protein is encoded by a transcript identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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determining a poorer prognosis if the level of the at least one protein is found to be higher in the first sample than in the second sample.

44. A method to aid in determining a prognosis of a patient having pancreatic cancer, comprising the steps of:

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comparing the level of at least one protein in a first sample to a second sample, wherein the first sample is of a pancreatic cancer patient and the second sample is of a normal human, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those shown Table 4, wherein said first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

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determining a poorer prognosis if the level of the at least one protein is found to be higher in the first sample than in the second sample.

45. A method to aid in providing a prognosis for a cancer patient, comprising the steps of:

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comparing the level of expression of at least one protein in a first sample to a second sample, wherein the first sample is of a cancer patient and the second sample is of a normal human, wherein said protein is encoded by a transcript identified by a tag selected from the group consisting of those

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shown Table 5, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one protein is found to be higher in the first sample than in the second sample.

A method of detecting colon cancer in a patient, comprising the steps of:

comparing the level of at least one transcript in a first body sample to a second body sample, wherein the first sample is a body sample of the patient and the second sample is of a normal human, wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

identifying neoplasia when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

47. A method of detecting pancreatic cancer in a patient, comprising the steps of:

comparing the level of at least one transcript in a first sample of a tissue to a second sample, wherein the first sample is of the patient and the second sample is of a normal human, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

identifying neoplasia when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

25 48. A method of detecting cancer in a patient, comprising the steps of:

comparing the level of at least one transcript in a first sample to
a second sample, wherein the first sample is of patient and the second sample
is of a normal human, wherein said transcript is identified by a tag selected

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from the group consisting of those shown Table 5, wherein the first and second body sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

identifying neoplasia when the level of the at least one transcript is found to be higher in the first sample than in the second sample.

49. A method to aid in determining a prognosis for a patient with colon cancer, comprising the steps of:

comparing the level of at least one transcript in a first sample to a second sample, wherein the first sample is of a colonic cancer patient and the second sample is of a normal human, wherein the transcript is identified by a tag selected from the group consisting of those shown in Table 2, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

50. A method to aid in determining a prognosis of a patient having pancreatic cancer, comprising the steps of:

comparing the level of at least one transcript in a first sample to a second sample, wherein the first sample is of a pancreatic cancer patient and the second sample is of a normal human, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 4, wherein said first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

51. A method to aid in providing a prognosis for a cancer patient, comprising the steps of:

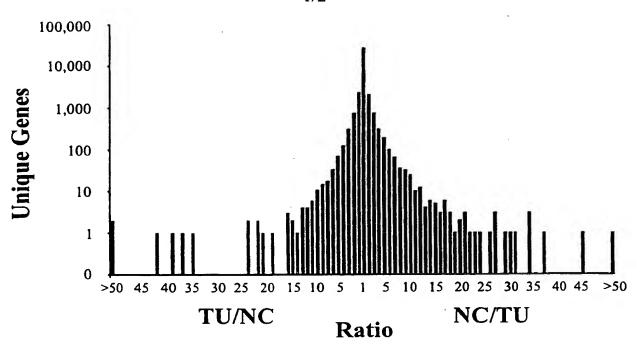
comparing the level of expression of at least one transcript in a first sample to a second sample, wherein the first sample is of a cancer patient and the second sample is of a normal human, wherein said transcript is identified by a tag selected from the group consisting of those shown Table 5, wherein the first and second sample is a sample selected from the group consisting of blood, urine, feces, sputum, and serum;

determining a poorer prognosis if the level of the at least one transcript is found to be higher in the first sample than in the second sample.

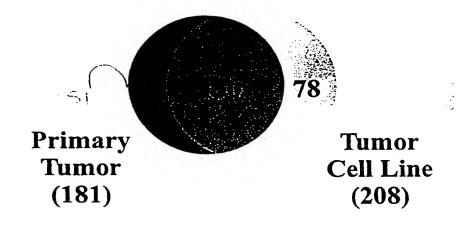
52. A method for screening for candidate agents that modulate the expression of a polynuleotide selected from the group consisting of the polynucleotides in SEQ ID NOS:1-732 or their respective complements, comprising contacting a test agent with a colon or pancreatic cell and monitoring expression of the polynucleotide, wherein the test agent which modifies the expression of the polynucleotide is a candidate agent.

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B.



C.

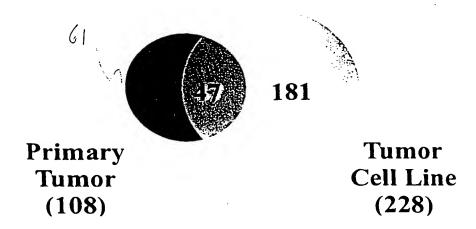


FIG. 2

A.

1 2 3	SAGE	Data
TNTNTN	T	N
H204104	.11	102
H259108	1	37
H1000193	56	12
H998030	55	7

B.

			_	ancre Tume					-	mal Ion	SAGE I)ata
	1	2	3	4	5	6	7	8	1	2	Pancreatic Tumors	Normai Colon
	H	-	H	H		-	1			H		
	4				Н					+1		
H294155	****	***		-	•	. •	44)		47	0
H560056		6				•	4)		32	0

C.

	CR Tumo	rs		ncrea umor:		Norr Colo		SA	GE Data	
	1 2	3	1	:		1 2			Pancreatic Tumors	Normal Colon
H802810								27	0	1
H85882			-				•	10	26	0
H618841			•	S	€,■	,		8	62	0

		.€ 	: :
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(71) Applicant (for all designated States except US): THE JOHNS HOPKINS UNIVERSITY [US/US]; Suite 2-100, 2024 E. Monument Street, Baltimore, MD 21205 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): VOGELSTEIN, Bert [US/US]; The Johns Hopkins University, Suite 2-100, 2024 E. Monument Street, Baltimore, MD 21205 (US). KIN-ZLER, Kenneth, W. [US/US]; The Johns Hopkins University, Suite 2-100, 2024 E. Monument Street, Baltimore, MD 21205 (US).

(74) Agents: KAGAN, Sarah, A. et al.; Banner & Witcoff, Ltd., 11th floor, 1001 G Street, N.W., Washington, DC 20001-4597

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(57) Abstract

As a step towards understanding the complex differences between normal and cancer cells, gene expression patterns were examined in gastrointestinal tumors. More than 300,000 transcripts derived from at least 45,000 different genes were analyzed. Although extensive similarity was noted between the expression profiles, more than 500 transcripts that were expressed at significantly different levels in normal and neoplastic cells were identified. These data provide insights into the extent of expression differences underlying malignancy and reveal genes that are useful as diagnostic or prognostic markers.

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A. CLASSIFICATION OF SUBJECT MATTER IPC 6 C12Q1/68 GOIN G01N33/574 According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) IPC 6 C12Q G01N Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Category ' Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. Х SUGIO K ET AL.: "Differential expression 1,3,13, of c-myc gene and c-fos gene in 16,17,28 premalignant and malignant tissues." CANCER RESEARCH, vol. 48, no. 17, 1988, pages 4855-4861, XP002089885 see the whole document 1,3,5,7, X VAN BELZEN N ET AL.: "Detection of 9.11 different gene expression in differentiating colon carcinoma cells by differential display" JOURNAL OF PATHOLOGY, vol. 178, no. Suppl., - 1996 page 2A XP002089886 Y see abstract 26,28,34 -/--Further documents are listed in the continuation of box C. Patent family members are listed in annex. Special categories of cited documents: "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the *A* document defining the general state of the lart which is not considered to be of particular meyance invention *E* earlier document but published on or after the international "X" document of particular relevance; the claimed invention filing date cannot be considered novel or cannot be considered to "L" document which may throw doubt on priority claim(s) or which is cited to establish the publication date of another involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention citation or other special reason (25 specified) cannot be considered to involve an inventive step when the document is combined with one or more other such docu-"O" document referring to an oral disclosure, use, exhibition or ments, such combination being obvious to a person skilled in the art. other means *P* document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report 2 4 05 1999 13 January 1999 Name and mailing address of the ISA Authorized officer European Patent Office, P. 3, 5o18 Patentiaan 2 NL - 2280 HV Risk F Tel. (+31-70) 340-20-1. Tr. 31 651 epo nl.

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Knehr, M

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Interi (nat Application No PCT/US 98/10277

ategory °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
(WO 95 21944 A (SMITHKLINE BEECHAM CORP;ROSENBERG MARTIN (US); DEBOUCK CHRISTINE) 17 August 1995 see the whole document	26,28,34
'	EP 0 284 362 A (ICI PLC) 28 September 1988	1,3,5,7, 9,11, 13-23, 26,28, 34,52
·	see abstract see page 2, line 44 - line 51 see page 10, line 12 - line 15; claims 1,9; figure 2	
Y	EP 0 761 822 A (UNIV JOHNS HOPKINS MED) 12 March 1997 see the whole document	1,3,5,7, 9,11, 13-23, 26,28, 34,52
	see the whole document	
Y	WO 95 11923 A (DANA FARBER CANCER INST INC ;CHEN LAN BO (US); BAO SHIDENG (CN); L) 4 May 1995	1,3,5,7, 9,11, 13-18, 23,26, 28,34,52
	see the whole document	20,34,32
Y	VELCULESCU V E ET AL: "SERIAL ANALYSIS OF GENE EXPRESSION" SCIENCE, vol. 270, 20 October 1995, pages 484-487, XP002053721 cited in the application see the whole document	1,3,5,7, 9,11, 13-18, 23,26, 28,34,52
Y	SCHWEINFEST C W ET AL.: "Subtraction hybridization cDNA libraries from colon carcinoma and hepatic cancer" GENETIC ANALYSIS TECHNIQUES AND APPLICATIONS, vol. 7, 1990, pages 64-70, XP002089887 see the whole document	1,3,5,7, 9,11, 13-18, 23,26
Υ	WO 97 14812 A (CHIRON CORP) 24 April 1997 see the whole document	52
Α	GRESS T M ET AL.: "A pancreatic cancer-specific expression profile" ONCOGENE, vol. 13, 1996, pages 1819-1830, XP002089888 see the whole document	

PCT/US 98/10277

ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No
WO 95 19369 A (UNIV VANDERBILT) 20 July 1995 see the whole document		
GRESS T ET AL.: "Identification of genes with pancreatic cancer specific expression by use of cDNA representational difference analysis" GASTROENTEROLOGY, vol. 110, no. 4 Suppl., 1996, XP002089889 see abstract		
ZHANG L E AL.: "Gene expression profiles in normal and cancer cells." SCIENCE, vol. 276, 1997, pages 1268-1272, XP002089890 see the whole document		1,3,5,7, 9,11, 13-23, 26,28, 34,52
VAN BELZEN N ET AL.: "A novel gene which is up-regulated during colon epithelial cell differentiation and down-regulated in colorectal neoplasms" LABORATORY INVESTIGATION, vol. 77, no. 1, 1997, pages 85-92, XP002089891 see the whole document		1,3,5,7, 9,11,13, 14, 16-18, 23,26, 28,34
	·	
	WO 95 19369 A (UNIV VANDERBILT) 20 July 1995 see the whole document GRESS T ET AL: "Identification of genes with pancreatic cancer specific expression by use of cDNA representational difference analysis" GASTROENTEROLOGY, vol. 110, no. 4 Suppl., 1996, XP002089889 see abstract ZHANG L E AL: "Gene expression profiles in normal and cancer cells." SCIENCE, vol. 276, 1997, pages 1268-1272, XP002089890 see the whole document VAN BELZEN N ET AL.: "A novel gene which is up-regulated during colon epithelial cell differentiation and down-regulated in colorectal neoplasms" LABORATORY INVESTIGATION, vol. 77, no. 1, 1997, pages 85-92, XP002089891 see the whole document	WO 95 19369 A (UNIV VANDERBILT) 20 July 1995 see the whole document GRESS T ET AL.: "Identification of genes with pancreatic cancer specific expression by use of cDNA representational difference analysis" GASTROENTEROLOGY, vol. 110, no. 4 Suppl., 1996, XP002089889 see abstract ZHANG L E AL.: "Gene expression profiles in normal and cancer cells." SCIENCE, vol. 276, 1997, pages 1268-1272, XP002089890 see the whole document VAN BELZEN N ET AL.: "A novel gene which is up-regulated during colon epithelial cell differentiation and down-regulated in colorectal neoplasms" LABORATORY INVESTIGATION, vol. 77, no. 1, 1997, pages 85-92, XP002089891 see the whole document

International application No.

PCT/US 98/10277

Box I	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This Inte	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1.	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
2.	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Inte	emational Searching Authority found multiple inventions in this international application, as follows:
se	e FURTHER INFORMATION sheet
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. X	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.: see FURTHER INFORMATION sheet, subject 1.
Remar	The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

International Application No. PCT/ US 98 / 10277

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FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: 1,3,5,7,9,11,13-23,26,28,34,52 (partial)

INVENTION 1:

An isolated and purified human nucleic acid molecule comprising SEQ ID NO:291 of table 3 (INVENTION 1), an isolated nucleotide probe hybridizing to it, a diagnostic reagent comprising at least two of such probes, methods of diagnosing or prognosing colon cancer using them, and a method for screening for agents modulating the expression of such a sequence using them.

2. Claims: 1,3,5,7,9,11,13-23,26,28,34,52 (partial)

INVENTION 2 to INVENTION 259:
An isolated and purified human nucleic acid molecule comprising SEQ ID NO:292 of table 3 (INVENTION 2), an isolated nucleotide probe hybridizing to it, a diagnostic reagent comprising at least two of such probes, methods of diagnosing or prognosing colon cancer using them, and a method for screening for agents modulating the expression of such a sequence using them.

...ibidem for each of the SEQ ID Nos:293 to 549 (INVENTION 3 to INVENTION 259) as specified in table 3, separately.

3. Claims: 2,4,6,8,10,12-23,27,29,35,38-40,43,46,49, 52 (partial)

INVENTION 260 to INVENTION 549:
An isolated and purified human nucleic acid molecule comprising SEQ ID NO:1 of table 2 (INVENTION 260), an isolated nucleotide probe hybridizing to it, a diagnostic reagent comprising at least two of such probes, methods of diagnosing or prognosing colon cancer using them, a method of treating a cancer cell using them, an antibody linked to a cytotoxic agent used in such a method, and a method for screening for agents modulating the expression of such a sequence using them.

...ibidem for each of the SEQ ID Nos:2 to 290 (INVENTION 261 to INVENTION 549) as specified in table 2, separately.

4. Claims: 13-24,30,32,36,38,39,41,44,47,50,52 (partial)

International Application No. PCT/FUS 98 / 10277

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

INVENTION 550 to INVENTION 732:
An isolated and purified human nucleic acid molecule comprising SEQ ID NO:550 of table 4 (INVENTION 550), an isolated nucleotide probe hybridizing to it, a diagnostic reagent comprising at least two of such probes, methods of diagnosing or prognosing pancreatic cancer using them, a method of treating a cancer cell using them, an antibody linked to a cytotoxic agent used in such a method, and a method for screening for agents modulating the expression of such a sequence using them.

...ibidem for each of the SEQ ID Nos:551 to 732 (INVENTION 551 to INVENTION 732) as specified in table 4, separately.

5. Claims: 24,30,32,36,38,39,41,44,47,50 (partial)

INVENTION 733 to INVENTION 734:
Methods of diagnosing or prognosing pancreatic cancer
relying on a human nucleic acid molecule comprising SEQ ID
NO:733 of table 4 (INVENTION 733), a method of treating a
cancer cell using it, and an antibody linked to a cytotoxic
agent used in such a method.

...ibidem for SEQ ID Nos:734 (INVENTION 734) as specified in table 4.

6. Claims: 25,31,33,37-39,42,45,48,51 (partial)

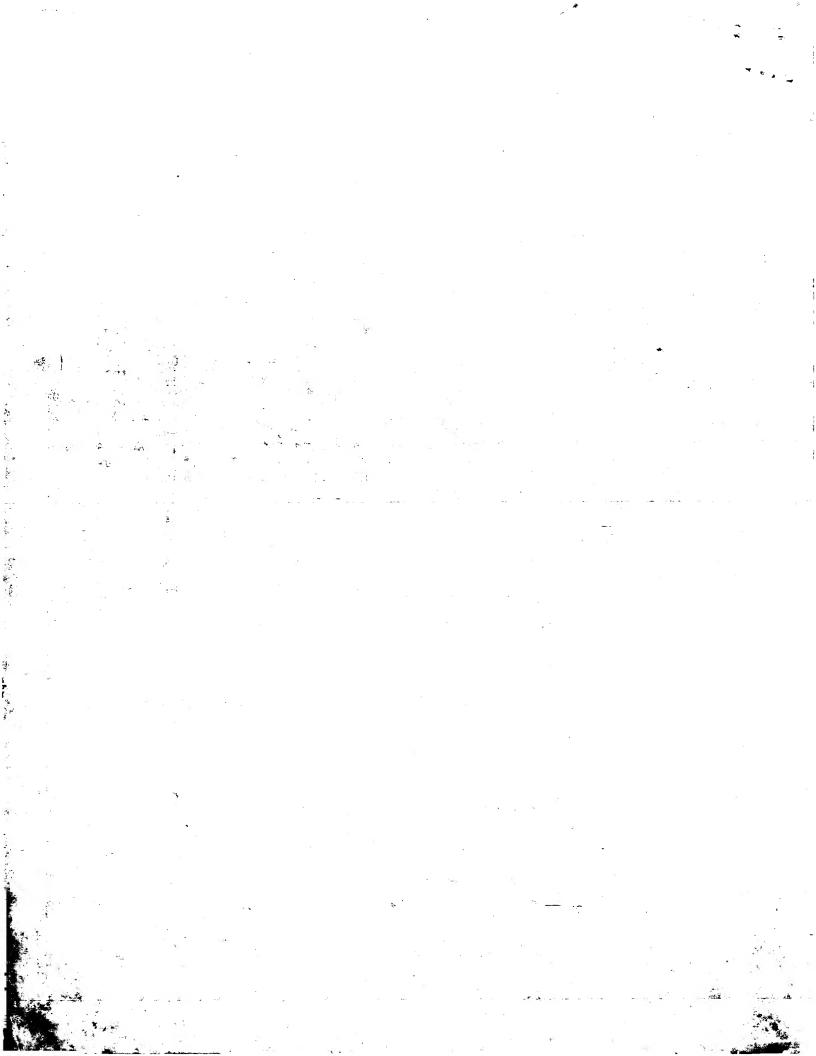
INVENTION 735 to INVENTION 870: Methods of diagnosing or prognosing cancer relying on a human nucleic acid molecule comprising SEQ ID NO:735 of table 5 (INVENTION 735), a method of treating a cancer cell using it, and an antibody linked to a cytotoxic agent used in such a method.

...ibidem for each of the SEQ ID Nos:736 to 870 (INVENTION 736 to INVENTION 870) as specified in table 5, separately.

.ormation on patent family members

Inte. .onal Application No PCT/US 98/10277

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
WO 9521944 A	17-08-1995	EP 0743989 A JP 9508800 T	27-11-1996 09-09-1997
EP 0284362 A	28-09-1988	AU 625169 B AU 1337888 A DK 159788 A FI 881388 A JP 1034291 A PT 87055 A,B	02-07-1992 -22-09-1988 -24-09-1988 -24-09-1988 -03-02-1989 -01-04-1988
EP 0761822 A	12-03-1997	US 5695937 A US 5866330 A AU 6561496 A AU 7018896 A CA 2185379 A GB 2305241 A IE 80465 B JP 10511002 T WO 9710363 A	09-12-1997 02-02-1999 20-03-1997 01-04-1997 13-03-1997 02-04-1997 12-08-1998 27-10-1998 20-03-1999
WO 9511923 A	04-05-1995	CA 2175380 A EP 0725799 A US 5889159 A US 5872235 A	04-05-1995 14-08-1996 30-03-1999 16-02-1999
WO 9714812 A	24-04-1997	AU 7264196 A EP 0862651 A	07-05-1997 09-09-1998
WO 9519369 A	20-07-1995	US 5677125 A AU 1831795 A CA 2210396 A EP 0804453 A	14-10-1997 01-08-1995 20-07-1995 05-11-1997



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(72) Inventors; and

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(54) Title: GENE EXPRESSION PROFILES IN NORMAL AND CANCER CELLS

(57) Abstract

As a step towards understanding the complex differences between normal and cancer cells, gene expression patterns were examined in gastrointestinal tumors. More than 300,000 transcripts derived from at least 45,000 different genes were analyzed. Although extensive similarity was noted between the expression profiles, more than 500 transcripts that were expressed at significantly different levels in normal and neoplastic cells were identified. These data provide insights into the extent of expression differences underlying malignancy and reveal genes that are useful as diagnostic or prognostic markers.

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A. CLASSIFICATION F SUBJECT MATTER
1PC 6 C12Q1/68 G01N33/574

According to international Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system tollowed by classification symbols) IPC 6 C12Q G01N

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

C. DOCUME	NTS CONSIDERED TO BE RELEVANT	
Category *	Citation of document, with indication,where appropriate, of the relevant passages	Relevant to claim No.
X	SUGIO K ET AL.: "Differential expression of c-myc gene and c-fos gene in premalignant and malignant tissues." CANCER RESEARCH, vol. 48, no. 17, 1988, pages 1881–1251, XP002089885 see the whole document	1,3,13, 16,17,28
X	VAN BELZEN N ET AL.: "Detection of different gene expression in differentiating colon carcinoma cells by differential display" JOURNAL OF PATHOLOGY, vol. 178, no. Suppl., - 1996	1,3,5,7,
Y	page 2A XP002089886 see abstract -/	26,28,34

Patent family members are listed in annex.		
T later document published after the international filing date or priority date and not in conflict with the application but cited to uncerstand the principle or theory underlying the invention.		
"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to		
"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the		
occurrent is combined with one or more other such docu- ments, such combination being obvious to a person skilled		
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C.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	. 以答為劉哲學學中
Category *	Chation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No
Y	WO 95 21944 A (SMITHKLINE BEECHAM CORP; ROSENBERG MARTIN (US); DEBOUCK CHRISTINE) 17 August 1995 see the whole document	26,28,34
Y	EP 0 284 362 A (ICI PLC) 28 September 1988	1,3,5,7, 9,11, 13-23, 26,28, 34,52
	see abstract see page 2, line 44 - line 51 see page 10, line 12 - line 15; claims 1,9; figure 2	
Y	EP 0 761 822 A (UNIV JOHNS HOPKINS MED) 12 March 1997	1,3,5,7, 9,11, 13-23,
	see the whole document	26.28, 34,52
Y	WO 95 11923 A (DANA FARBER CANCER INST INC ;CHEN LAN BO (US); BAO SHIDENG (CN); L) 4 May 1995	1,3,5,7, 9,11, 13-18, 23,26, 28,34,52
Y	velculescu v e et al: "Serial analysis of GENE EXPRESSION" SCIENCE, vol. 270, 20 October 1995, pages 484-487, XP002053721 cited in the application see the whole document	1,3,5,7, 9,11, 13-18, 23,26, 28,34,52
Y	SCHWEINFEST C W ET AL.: "Subtraction hybridization cDNA libraries from colon carcinoma and hepatic cancer" GENETIC ANALYSIS TECHNIQUES AND APPLICATIONS, vol. 7, 1990, pages 64-70, XP002089887 see the whole document	1,3,5,7, 9,11, 13-18, 23,26
Υ	WO 97 14812 A (CHIRON CORP) 24 April 1997 see the whole document	52
A	GRESS T M ET AL.: "A pancreatic cancer-specific expression profile" ONCOGENE, vol13, 1996,	
	pages 1819-1830, XP002089888 see the whole document	

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C /Continua	tion)-DOCUMENTS CONSIDERED TO BE RELEVANT	J	
Category *	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No
A	WO 95 19369 A (UNIV VANDERBILT) 20 July 1995 see the whole document		
A .	GRESS T ET AL.: "Identification of genes with pancreatic cancer specific expression by use of cDNA representational difference analysis" GASTROENTEROLOGY, vol. 110, no. 4 Suppl., 1996, XP002089889 see abstract		
P,X	ZHANG L E AL.: "Gene expression profiles in normal and cancer cells." SCIENCE, vol. 276, 1997, pages 1268-1272, XP002089890 see the whole document		1,3,5,7, 9,11, 13-23, 26,28, 34,52
P;X	VAN BELZEN N ET AL.: "A novel gene which is up-regulated during colon epithelial cell differentiation and down-regulated in colorectal neoplasms" LASOKATOR: INVESTIGATION, vol. 77, no. 1, 1997, pages 85-92, XP002089891 see the whole document		1,3,5,7, 9,11,13, 14, 16-18, 23,26, 28,34
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FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: 1,3,5,7,9,11,13-23,26,28,34,52 (partial)

INVENTION 1:

An isolated and purified human nucleic acid molecule comprising SEQ ID NO:291 of table 3 (INVENTION 1), an isolated nucleotide probe hybridizing to it, a diagnostic reagent comprising at least two of such probes, methods of diagnosing or prognosing colon cancer using them, and a method for screening for agents modulating the expression of such a sequence using them.

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2. Claims: 1,3,5,7,9,11,13-23,26,28,34,52 (partial)

INVENTION 2 to INVENTION 259:
An isolated and purified human nucleic acid molecule comprising SEQ ID NO:292 of table 3 (INVENTION 2), an isolated nucleotide probe hybridizing to it, a diagnostic reagent comprising at least two of such probes, methods of diagnosing or prognosing colon cancer using them, and a method for screening for agents modulating the expression of such a sequence using them.

- ...ibidem for each of the SEQ ID Nos:293 to 549 (INVENTION 3 to INVENTION 259) as specified in table 3, separately.
- 3. Claims: 2,4,6,8,10,12-23,27,29,35,38-40,43,46,49, 52 (partial)

INVENTION 260 to INVENTION 549:
An isolated and purified human nucleic acid molecule comprising SEQ ID NO:1 of table 2 (INVENTION 260), an isolated nucleotide probe hybridizing to it, a diagnostic reagent comprising at least two of such probes, methods of diagnosing or prognosing colon cancer using them, a method of treating a cancer cell using them, an antibody linked to a cytotoxic agent used in such a method, and a method for screening for agents modulating the expression of such a sequence using them.

- ...ibidem for each of the SEQ ID Nos:2 to 290 (INVENTION 261 to INVENTION 549) as specified in table 2, separately.
- 4. Claims: 13-24,30,32,36,38,39,41,44,47,50,52 (partial)

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International Application No. PCT/US 98 / 10277

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

INVENTION 550 to INVENTION 732:
An isolated and purified human nucleic acid molecule comprising SEQ ID NO:550 of table 4 (INVENTION 550), an isolated nucleotide probe hybridizing to it, a diagnostic reagent comprising at least two of such probes, methods of diagnosing or prognosing pancreatic cancer using them, a method of treating a cancer cell using them, an antibody linked to a cytotoxic agent used in such a method, and a method for screening for agents modulating the expression of such a sequence using them.

...ibidem for each of the SEQ ID Nos:551 to 732 (INVENTION 551 to INVENTION 732) as specified in table 4, separately.

5. Claims: -24,30,32,36,38,39,41,44,47,50 (partial)

INVENTION 733 to INVENTION 734:
Methods of diagnosing or prognosing pancreatic cancer
relying on a human nucleic acid molecule comprising SEQ ID
NO:733 of table 4 (INVENTION 733), a method of treating a
cancer cell using it, and an antibody linked to a cytotoxic
agent used in such a method.

...ibidem for SEQ ID Nos:734 (INVENTION 734) as specified in table 4.

6. Claims: 25,31,33,37-39,42,45,48,51 (partial)

INVENTION 735 to INVENTION 870:
Methods of diagnosing or prognosing cancer relying on a
human nucleic acid molecule comprising SEQ ID NO:735 of
table 5 (INVENTION 735), a method of treating a cancer cell
using it, and an antibody linked to a cytotoxic agent used
in such a method.

...ibidem for each of the SEQ ID Nos:736 to 870 (INVENTION 736 to INVENTION 870) as specified in table 5, separately.

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	Patent document cited in search report	rt	Publication date	Patent family member(s)		Publication date	
	WO 9521944	A	17-08-1995	EP 07439 JP 95088		27-11-1996 09 - 09-1997	·
	EP 0284362	A	28-09-1988	AU 6251 AU 13378 DK 1597 FI 8813 JP 10342 PT 870	38 A 38 A 38 A	02-07-1992 22-09-1988 24-09-1988 24-09-1988 03-02-1989 01-04-1988	
	EP 0761822	А	12-03-1997	US 56959 US 58663 AU 65614 AU 70188 CA 21853 GB 230524 IE 8044 JP 105110 WO 97103	30 A 96 A 96 A 79 A 41 A 55 B	09-12-1997 02-02-1999 20-03-1997 01-04-1997 13-03-1997 02-04-1997 12-08-1998 27-10-1998 20-03-1999	
	w0 9511923	A	04-05-1995	CA 21753 EP 07257 US 58891 US 58722	99 A 59 A	04-05-1995 14-08-1996 30-03-1999 16-02-1999	
	WO 9714812	A	24-04-1997	AU 72641 EP 08626		07-05-1997 09-09-1998	
	WO 9519369	A	20-07-1995	US 56771 AU 18317 CA 22103 EP 08044	95 A 96 A	14-10-1997 01-08-1995 20-07-1995 05-11-1997	
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